A person in silhouette is running on a beach at sunset. The person is in the foreground, moving from right to left. The background shows the ocean and a colorful sky with orange, red, and blue hues. The text is overlaid on the left side of the image.

*“Natural science, does not simply describe and explain nature; it is part of the interplay between nature and ourselves.”*

*– Werner Heisenberg*

**Module II: Computations in the Biological World, Lecture III.b**

**Chi-Ning Chou @ 2022 January Mini-Course “What is Computation? From Turing Machines to Black Holes and Neurons”**



# Two Mysteries: Evolution and Neuroscience

## Module III: Computations in the Biological World

*“Natural science, does not simply describe and explain nature; it is part of the interplay between nature and ourselves.”*

*– Werner Heisenberg*

# Last Lecture

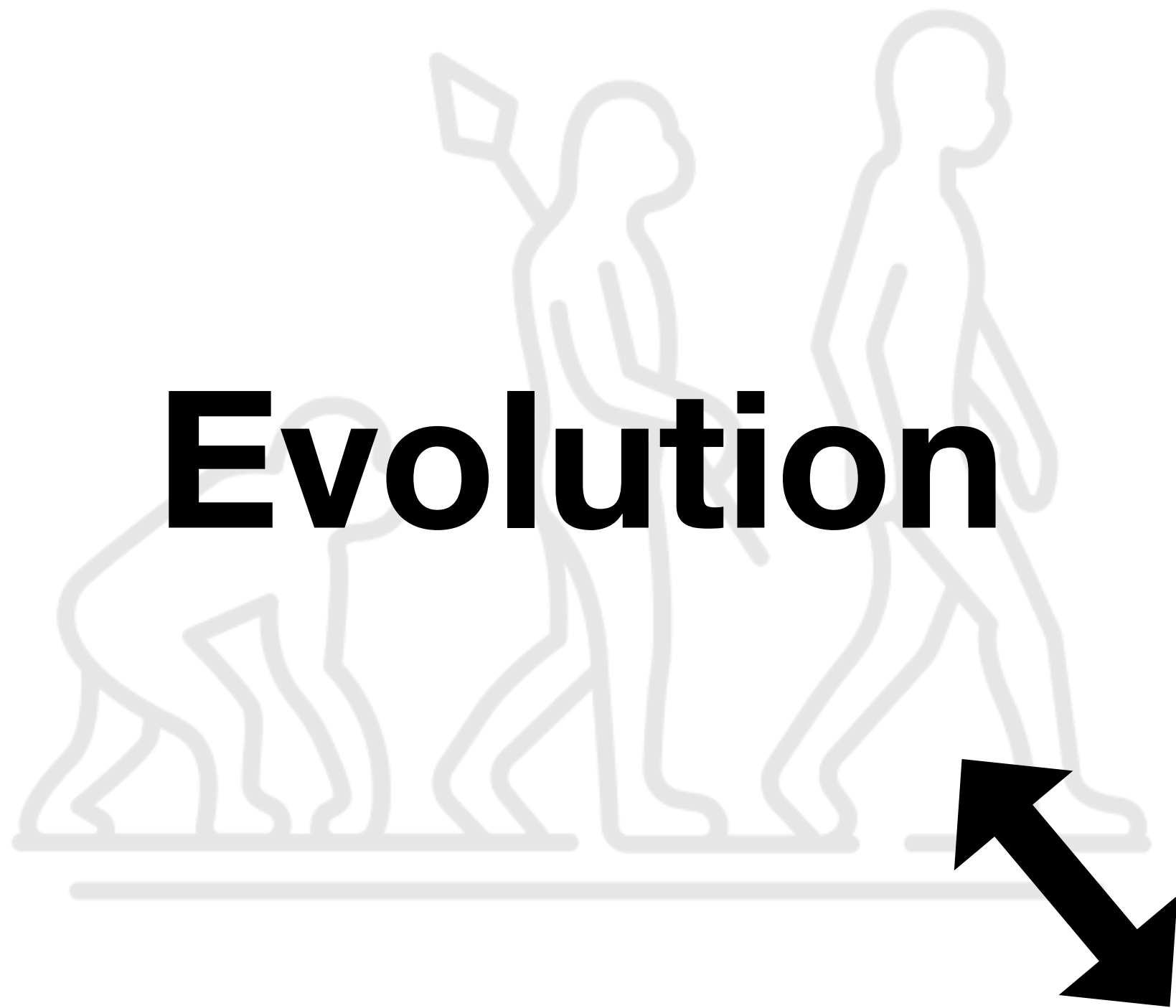
- What is biology?
- Marr's three levels.
- Computational biology & bioinformatics.
- Biological computations.

- Evolution.
- Neuroscience.

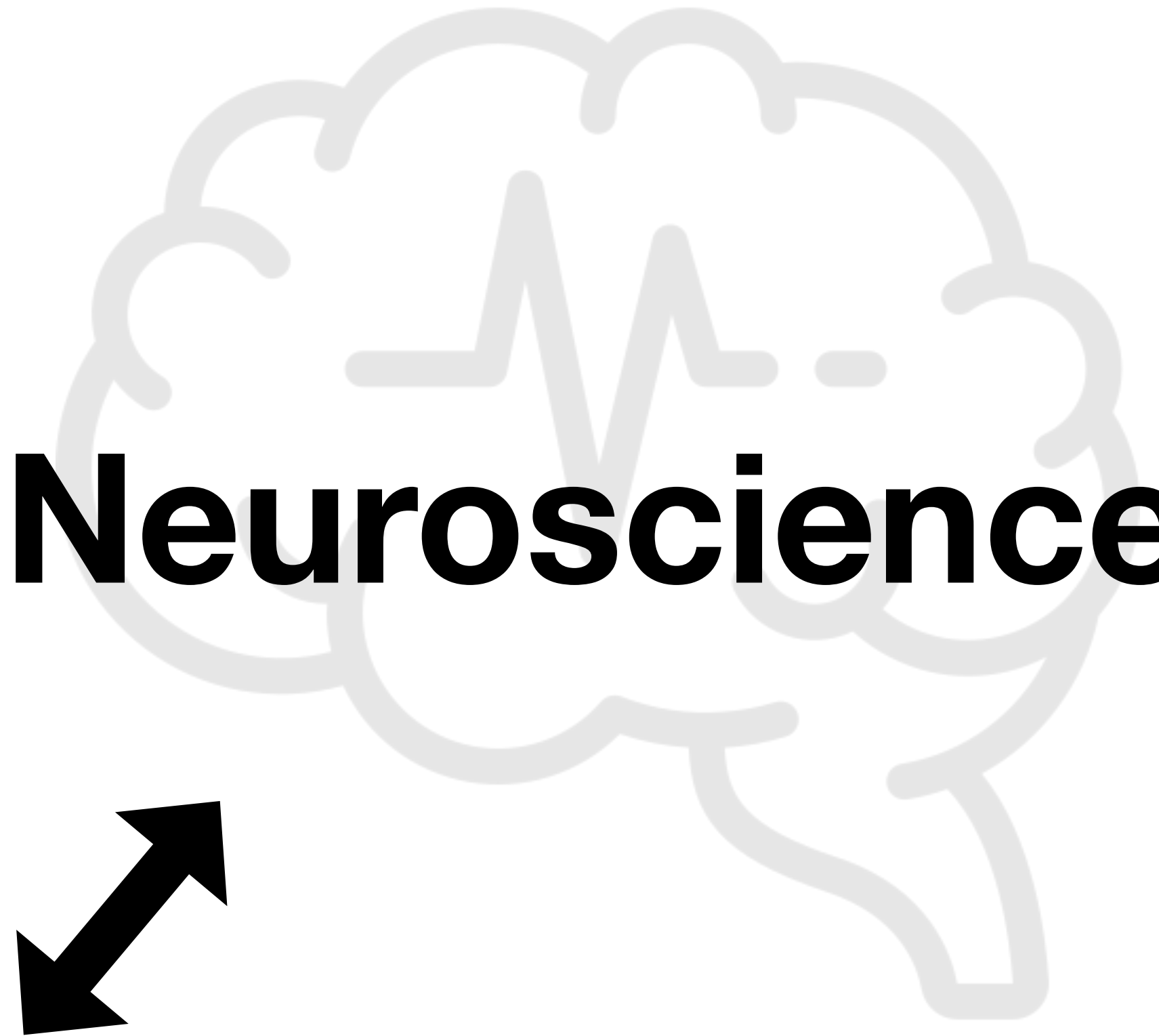
## **This Lecture**

# Two Mysteries

**Evolution**



**Neuroscience**



**Computation**



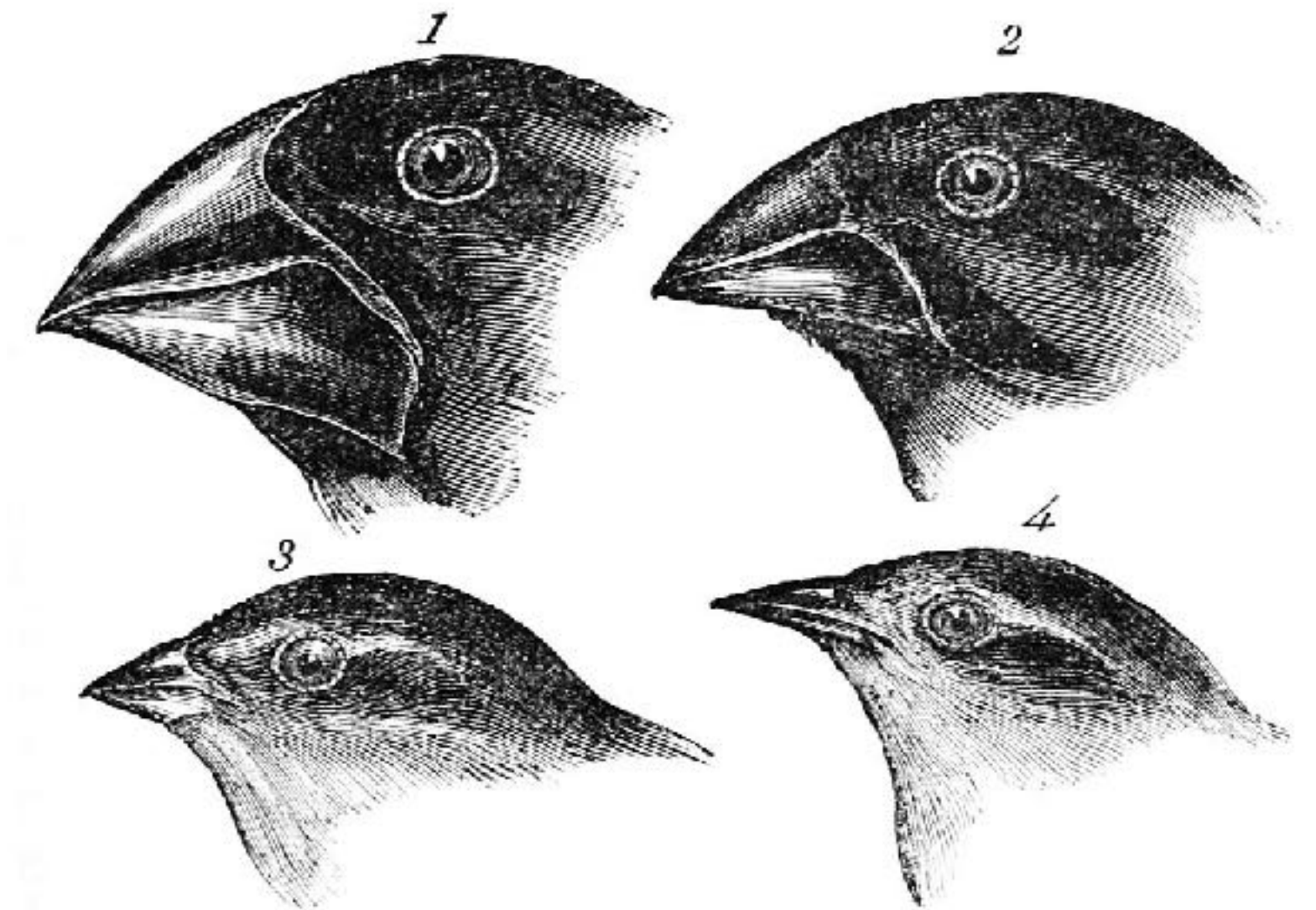
# Evolution

*“Evolution has no long-term goal. There is no long-distance target, no final perfection to serve as a criterion for selection.”*

*– Richard Dawkins*



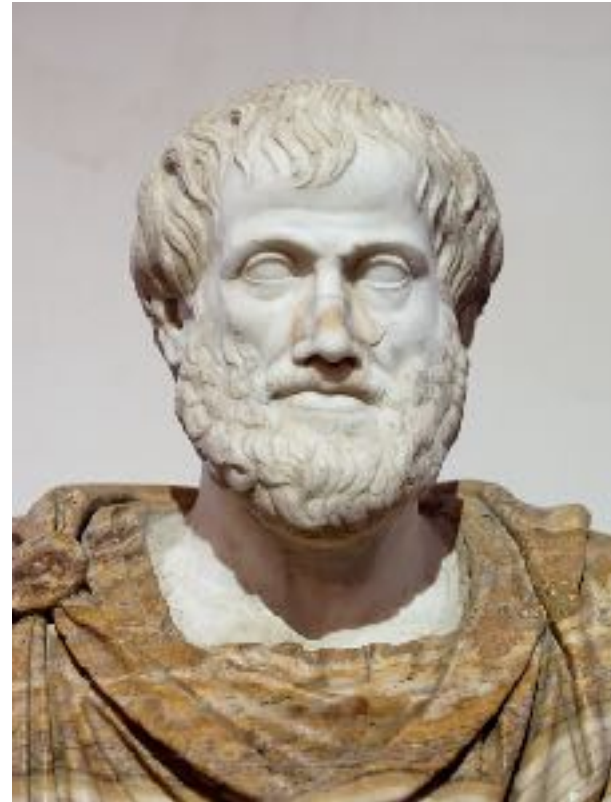
# Structure and Randomness



What's the underlying reason for diversity and similarity?



# Before and After Darwin's Journey

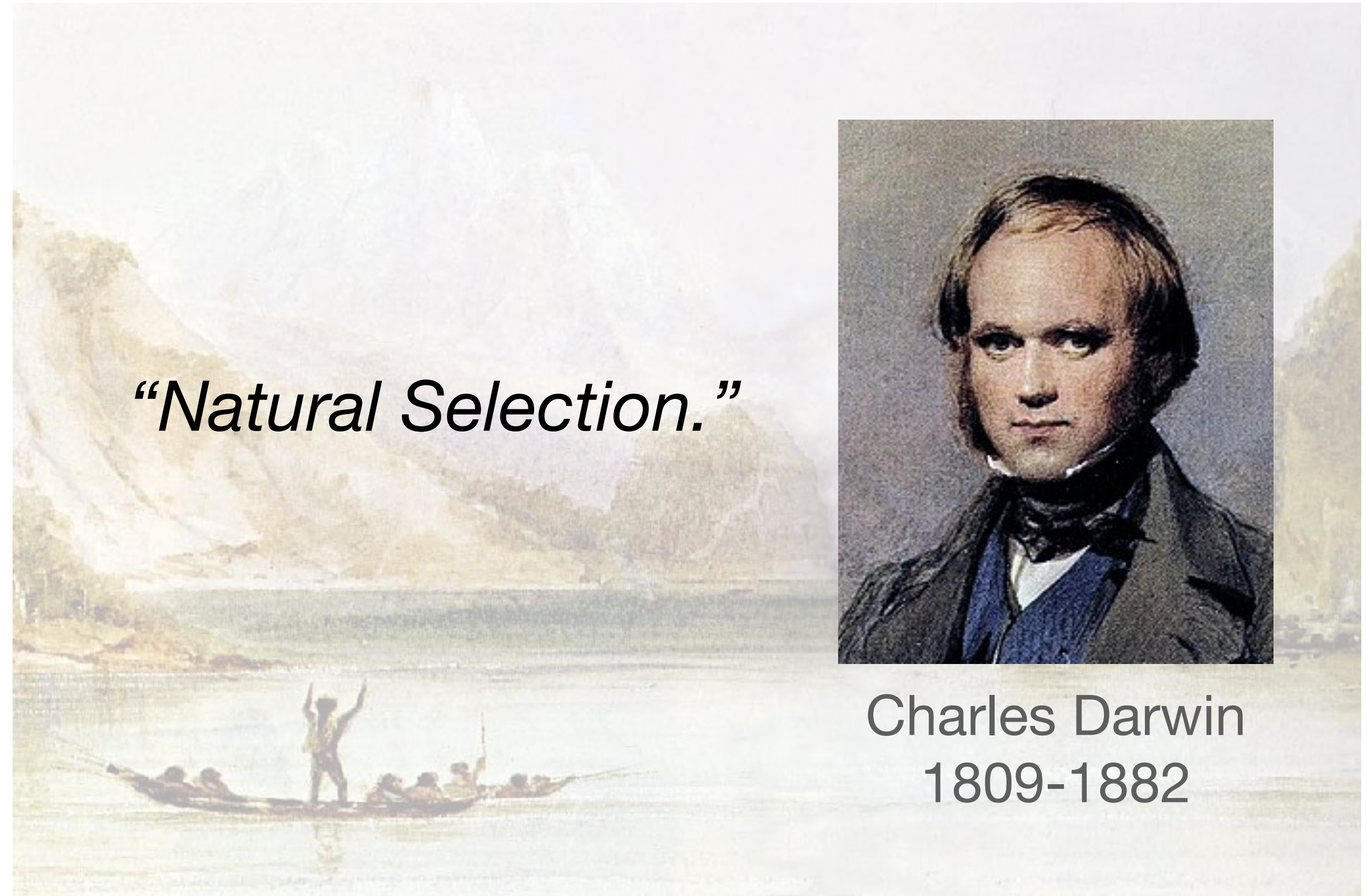


Aristotle  
384BC-322BC

*“The acquired  
characteristics  
can be inherited.”*



Jean-Baptiste Lamarck  
1744-1829



*“Natural Selection.”*



Charles Darwin  
1809-1882

The Beagle Expedition  
1831-1836



ON  
THE ORIGIN OF SPECIES  
BY MEANS OF NATURAL SELECTION,  
OR THE  
PRESERVATION OF FAVOURED RACES IN THE STRUGGLE  
FOR LIFE.

By CHARLES DARWIN, M.A.,  
FELLOW OF THE ROYAL, GEOLOGICAL, LINNÆAN, ETC., SOCIETIES;  
AUTHOR OF 'JOURNAL OF RESEARCHES DURING H. M. S. BEAGLE'S VOYAGE  
ROUND THE WORLD.'

LONDON:  
JOHN MURRAY, ALBEMARLE STREET.  
1859.

*The right of Translation is reserved.*

# The Origin of Species

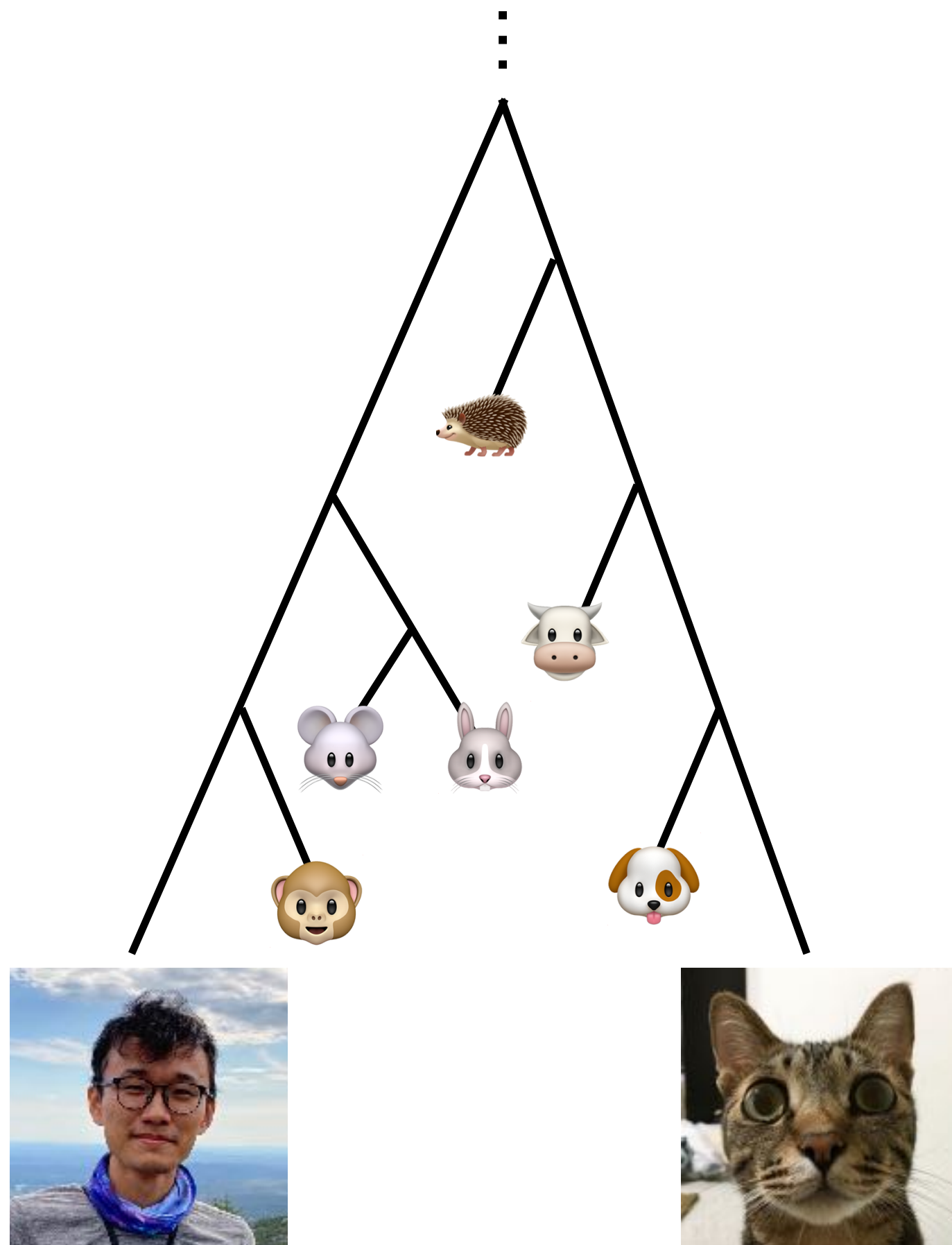
## By Means of Natural Selection,

or the

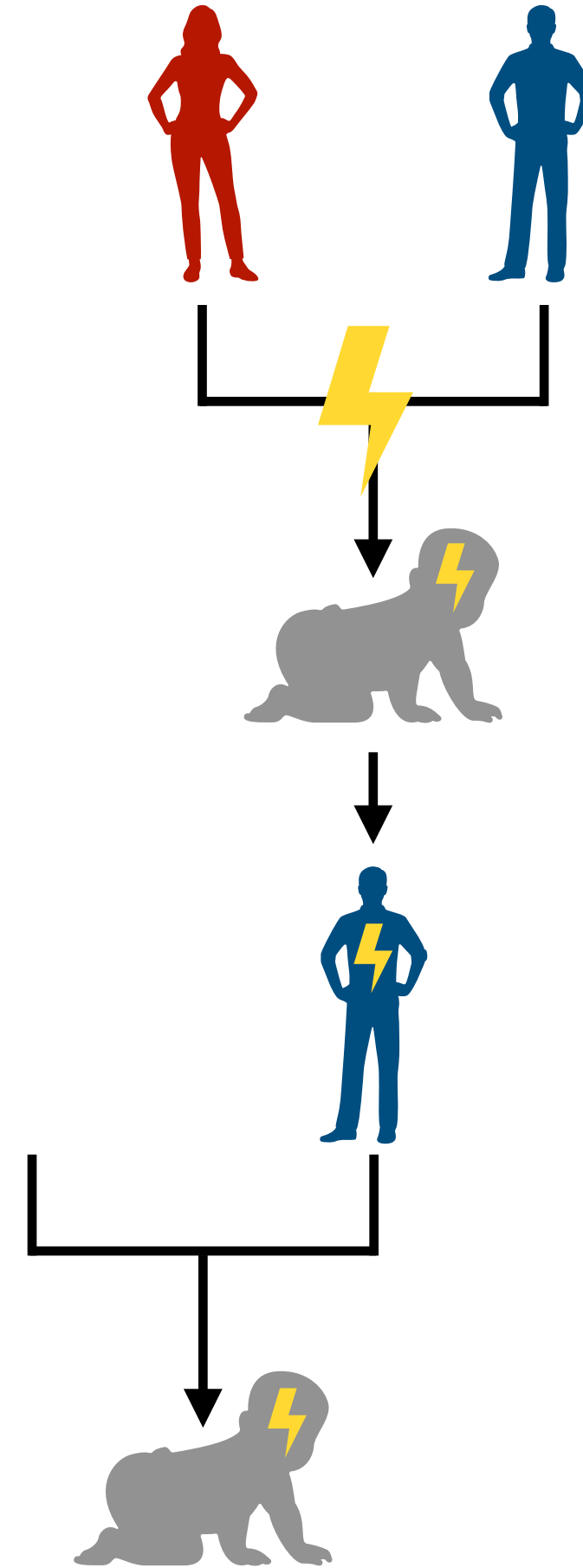
## Preservation of Favoured Races in the Struggle for Life.



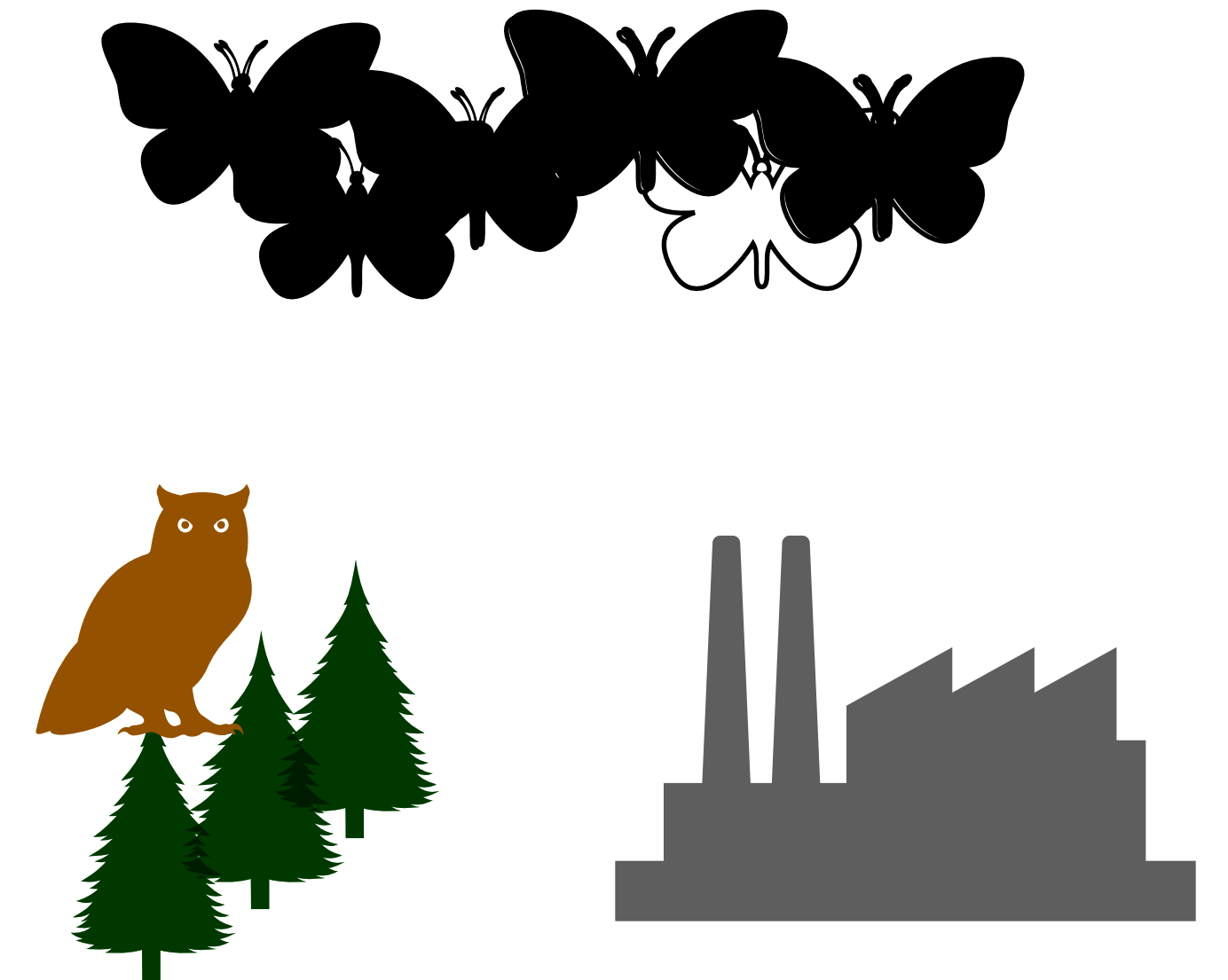
# The Essence of Natural Selection



**Common descent**



**Heredity & Variation**



**Competition & Fitness**



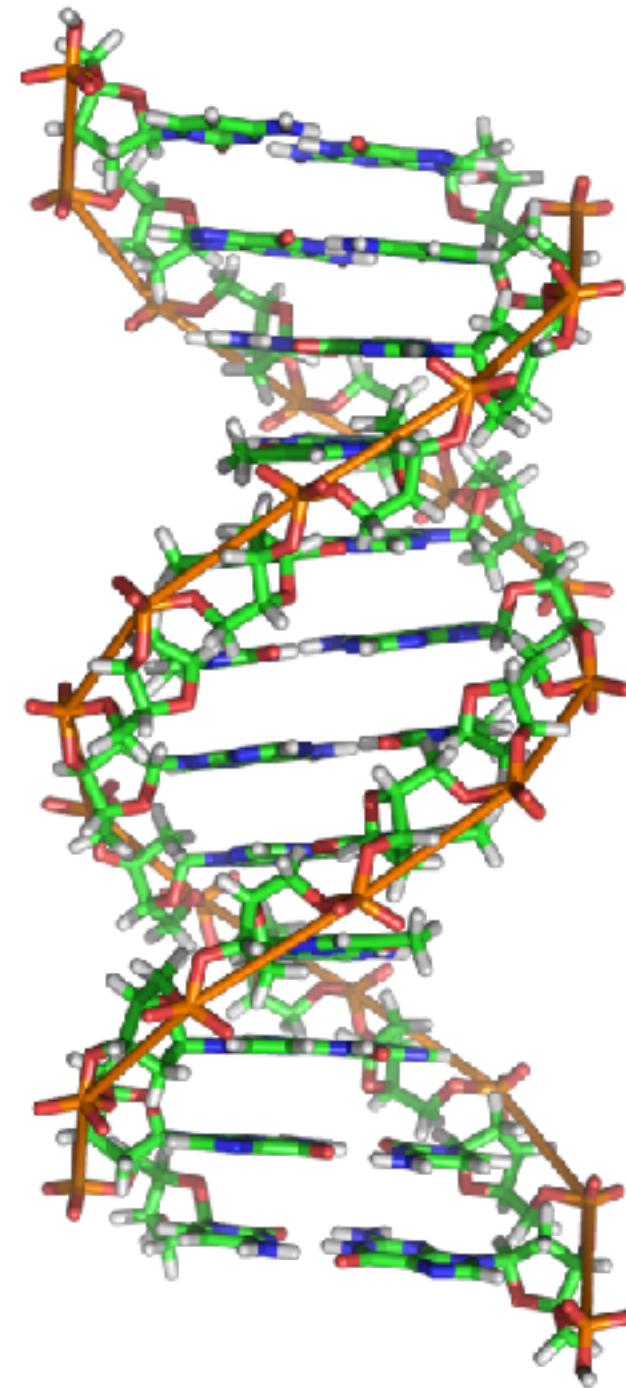
# How to “Prove” the Validity of Natural Selection?

*“Well, evolution is a theory. It is also a fact. And facts and theories are different things, not rungs in a hierarchy of increasing certainty. Facts are the world’s data. Theories are structures of ideas that explain and interpret facts.”*

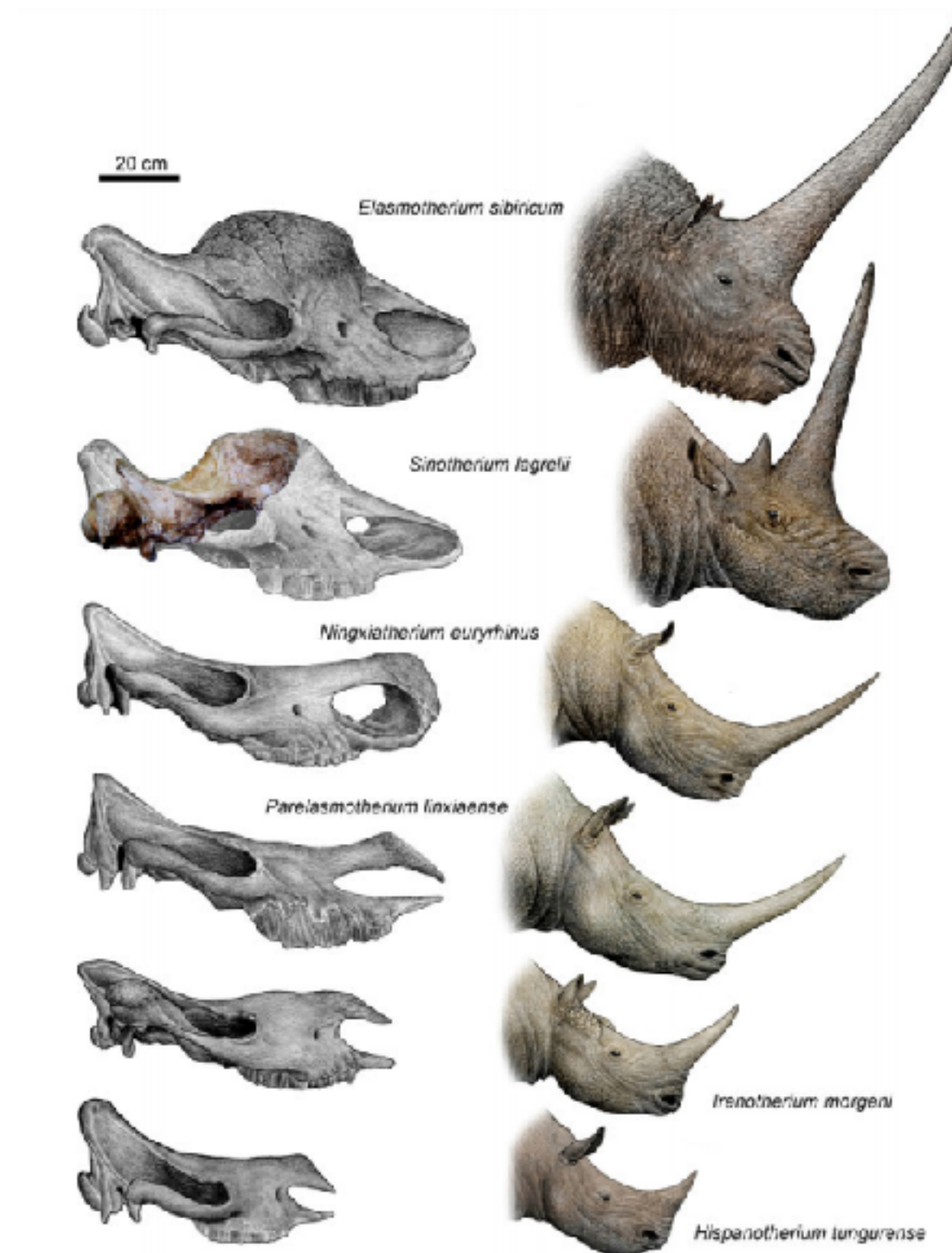
*– Stephen Jay Gould*



# Evidence Supporting Natural Selection



Genetics



Fossils

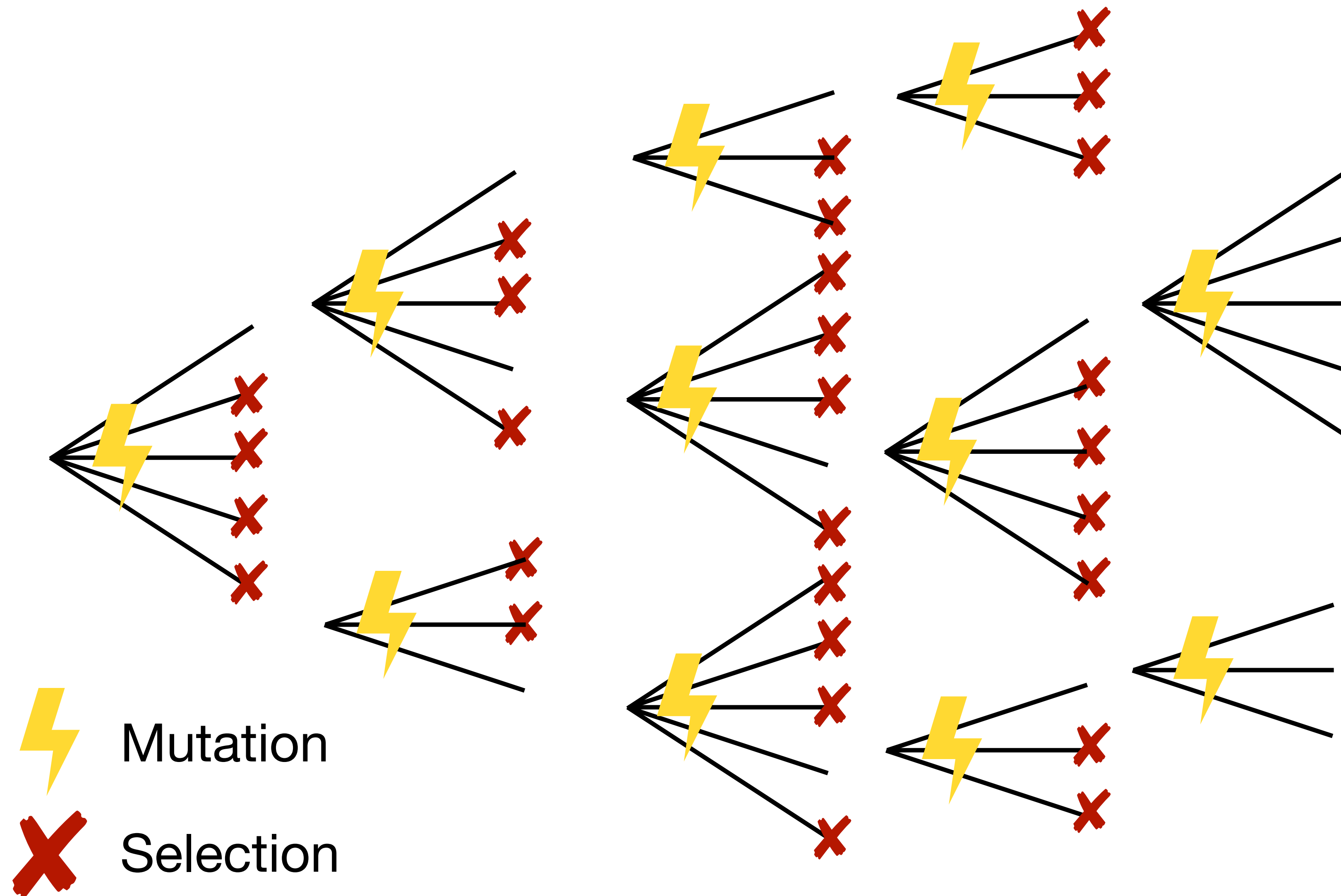


Lecture II.c

(Jan. 20 10am-10:50am ET)



# A New Angle to See the World



**Q:** What's the underlying computation!?

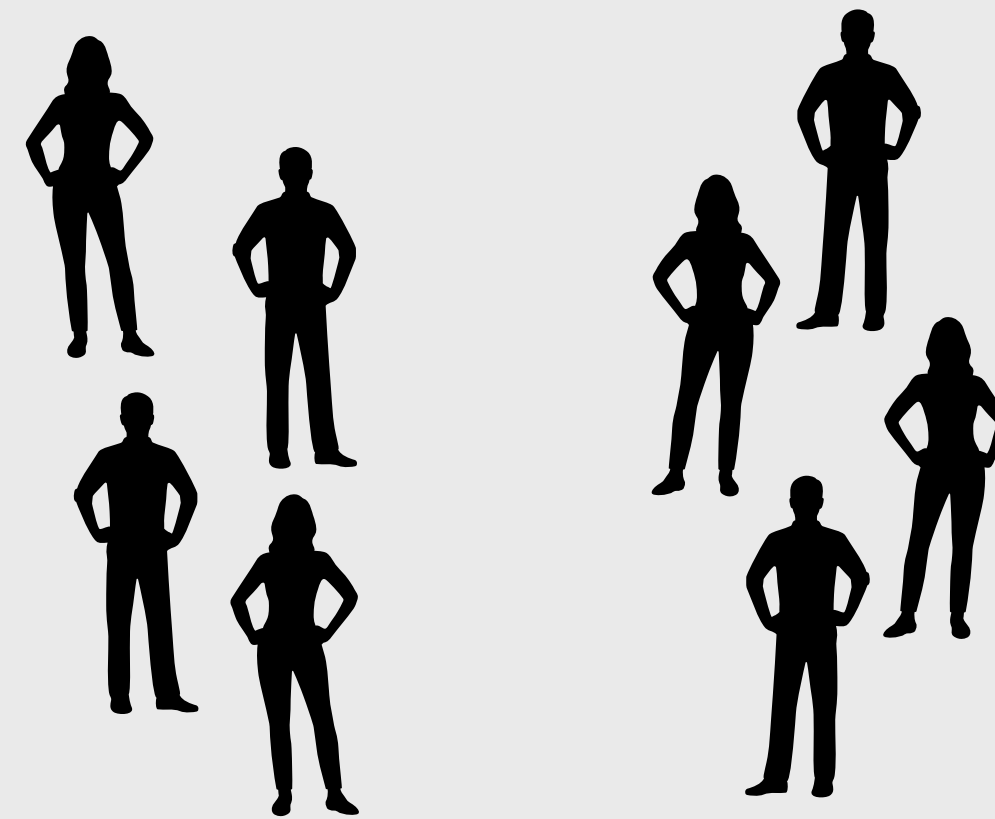


# Examples of Evolutionary Thinking

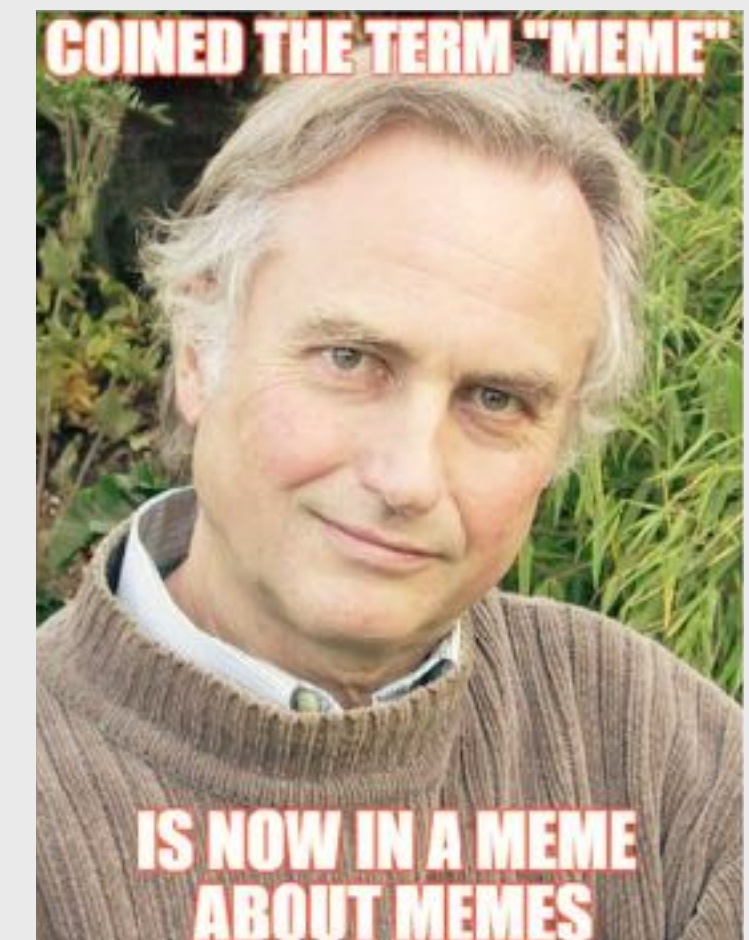
## Example 1: Sexual Dimorphism



## Example 2: Group Selection

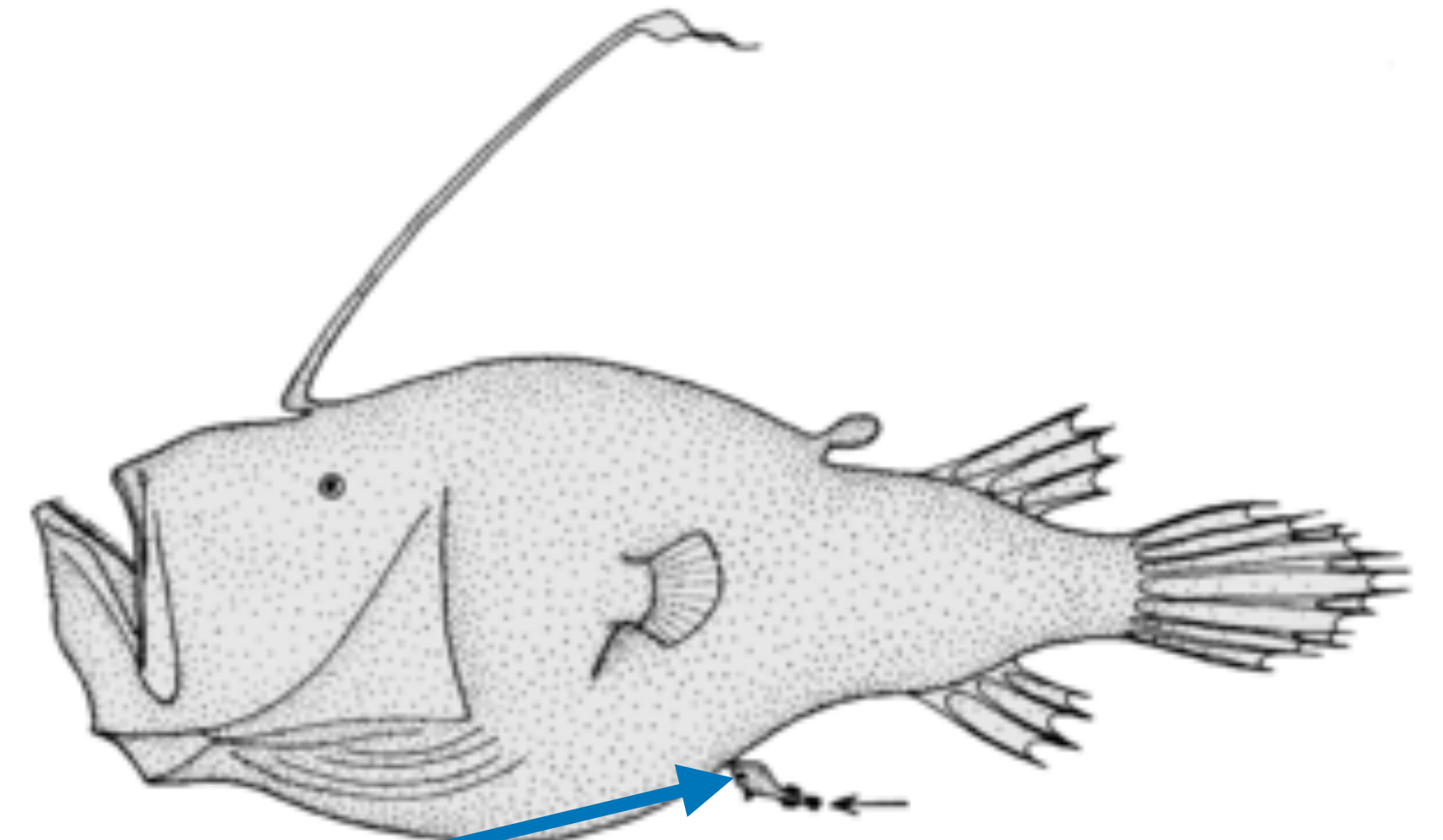
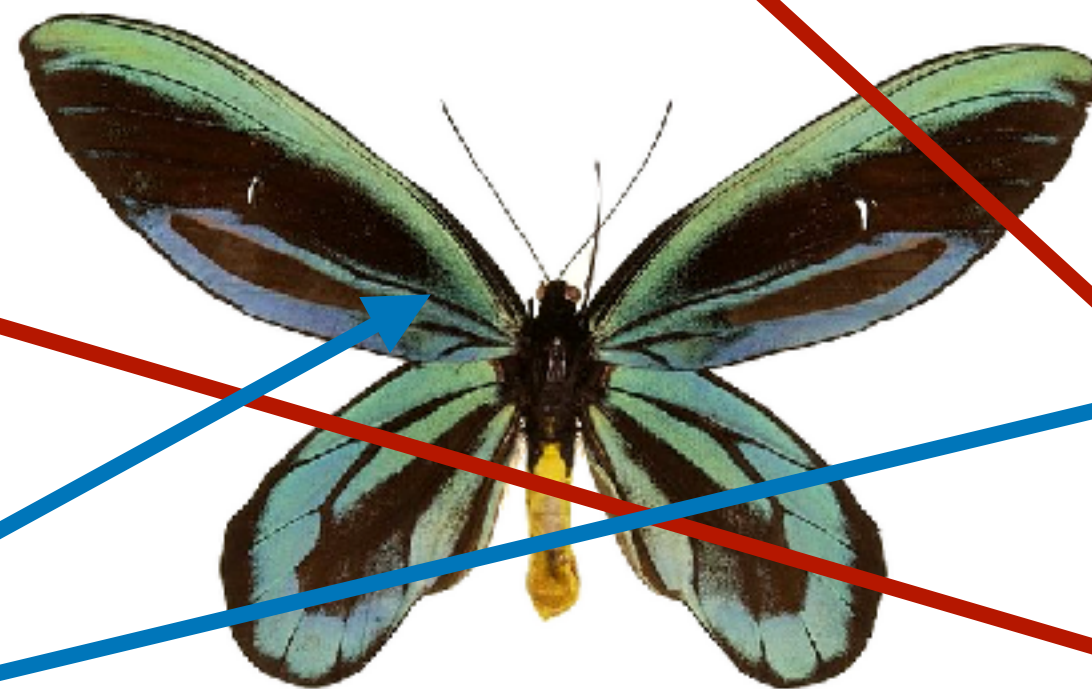
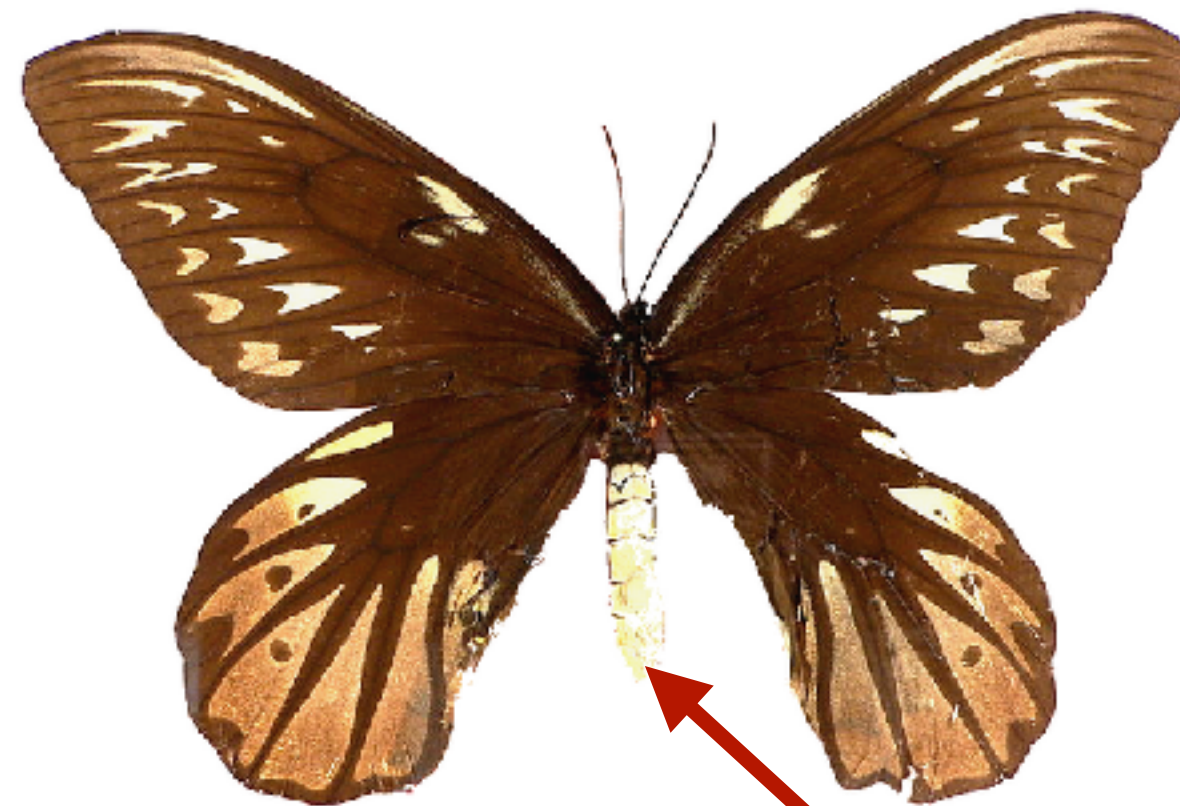


## Example 3: Meme





# Example 1: Sexual Dimorphism



**Male**

**Female**



# Two Types of Sexual Selection

*There are other new theories proposed in modern studies!*

## Intersexual selection



## Intrasexual selection



A mixture of these two selection forces?

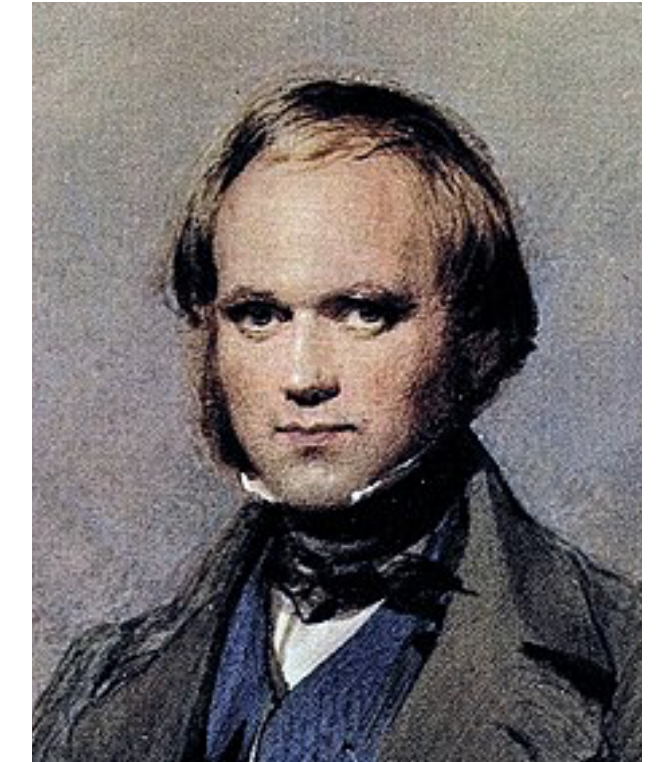


# Example 2: Group Selection

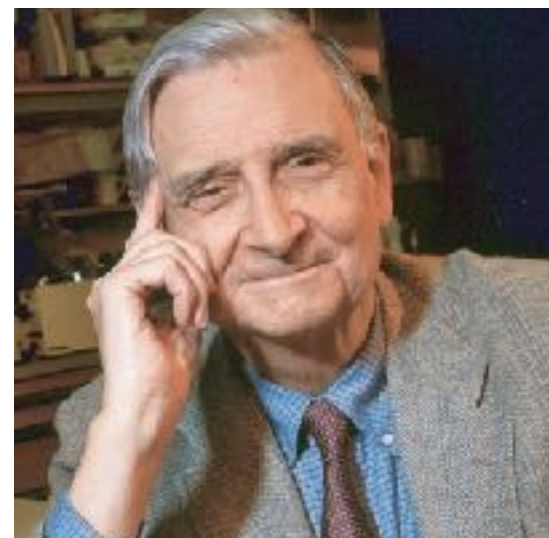
*“A tribe of moral men have an immense advantage over a group of fractious bands of pirates!?”*

Intuitively, this seems to make sense, but how to scientifically argue it? Difficulties are:

- Selection happens in multilevel.
- Selection forces at different levels might contradict to each other.



THE  
DESCENT OF MAN,



E. O. Wilson  
1921-2021

*“In a group, selfish individuals beat altruistic individuals. But, groups of altruistic individuals beat groups of selfish individuals.”*

*– E. O. Wilson*



# The Foundress's Dilemma



**Aggressive**

Lower colony density

Both types of queen exist!  
Why!?



**Cooperative**

Higher colony density



In other cases of group selection, there could be more complicated  
structure of selection forces!



# Example 3: Meme



Richard Dawkins  
1941-present

*“We need a name for the new replicator,  
a noun that conveys the idea of a unit  
of cultural transmission, or a unit or  
imitation.”*

*– Richard Dawkins*

More than the memes on the internet...



**Politics**



**Religion**



**Science**

...





# Examples of Evolutionary Thinking

## Example 1:

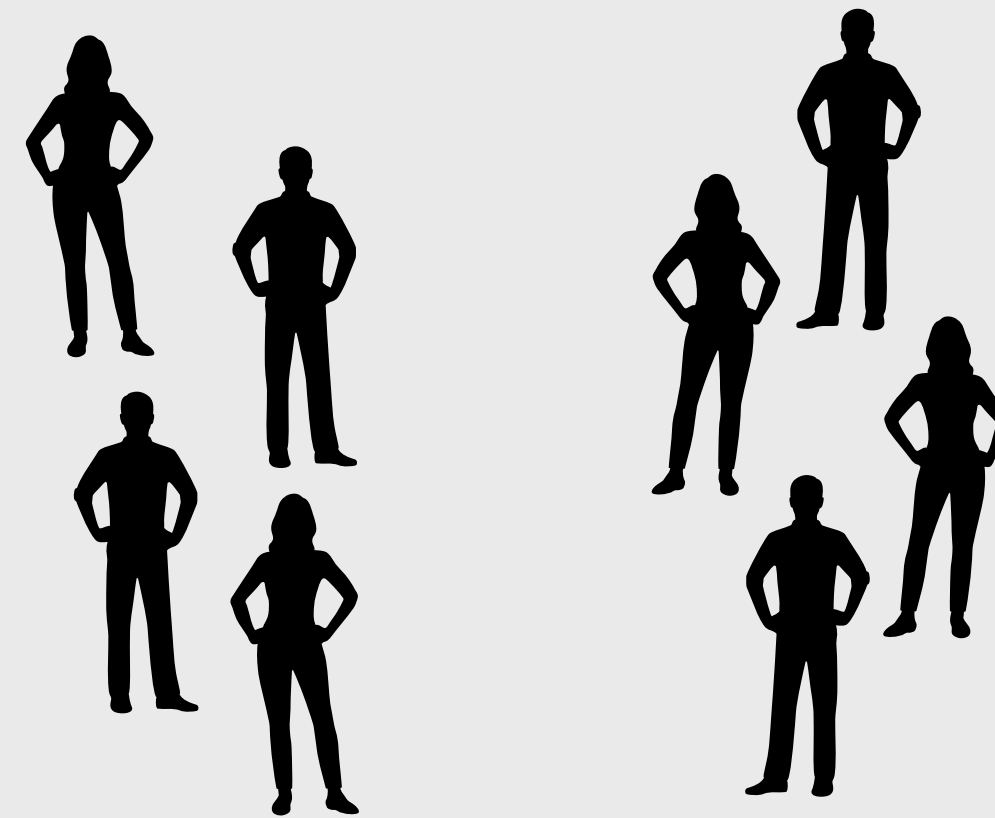
Sexual Dimorphism



Multiple  
selection forces

## Example 2:

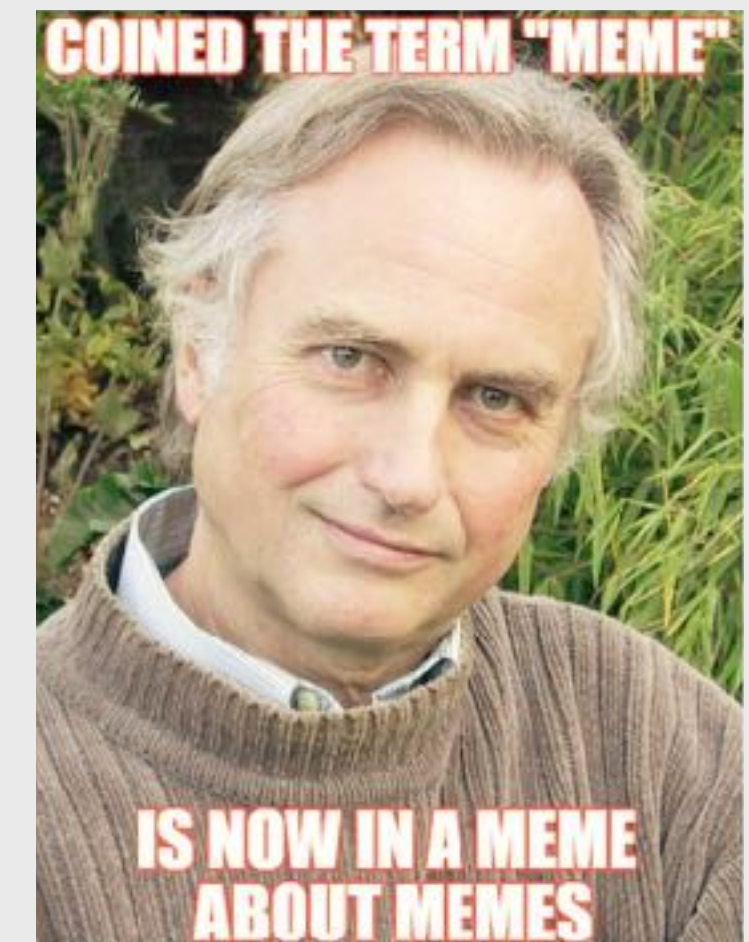
Group Selection



Multiple  
levels of selection

## Example 3:

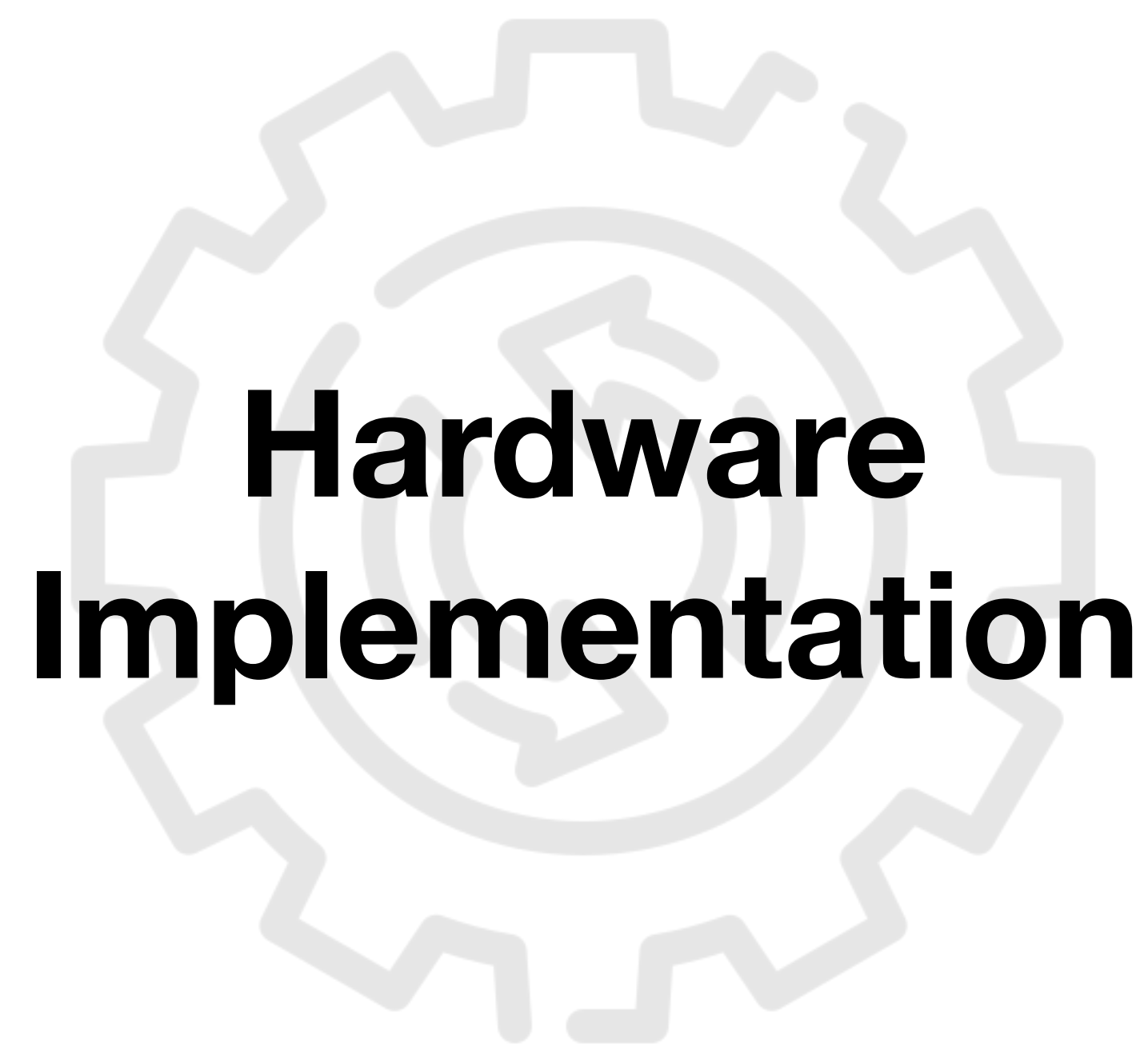
Meme



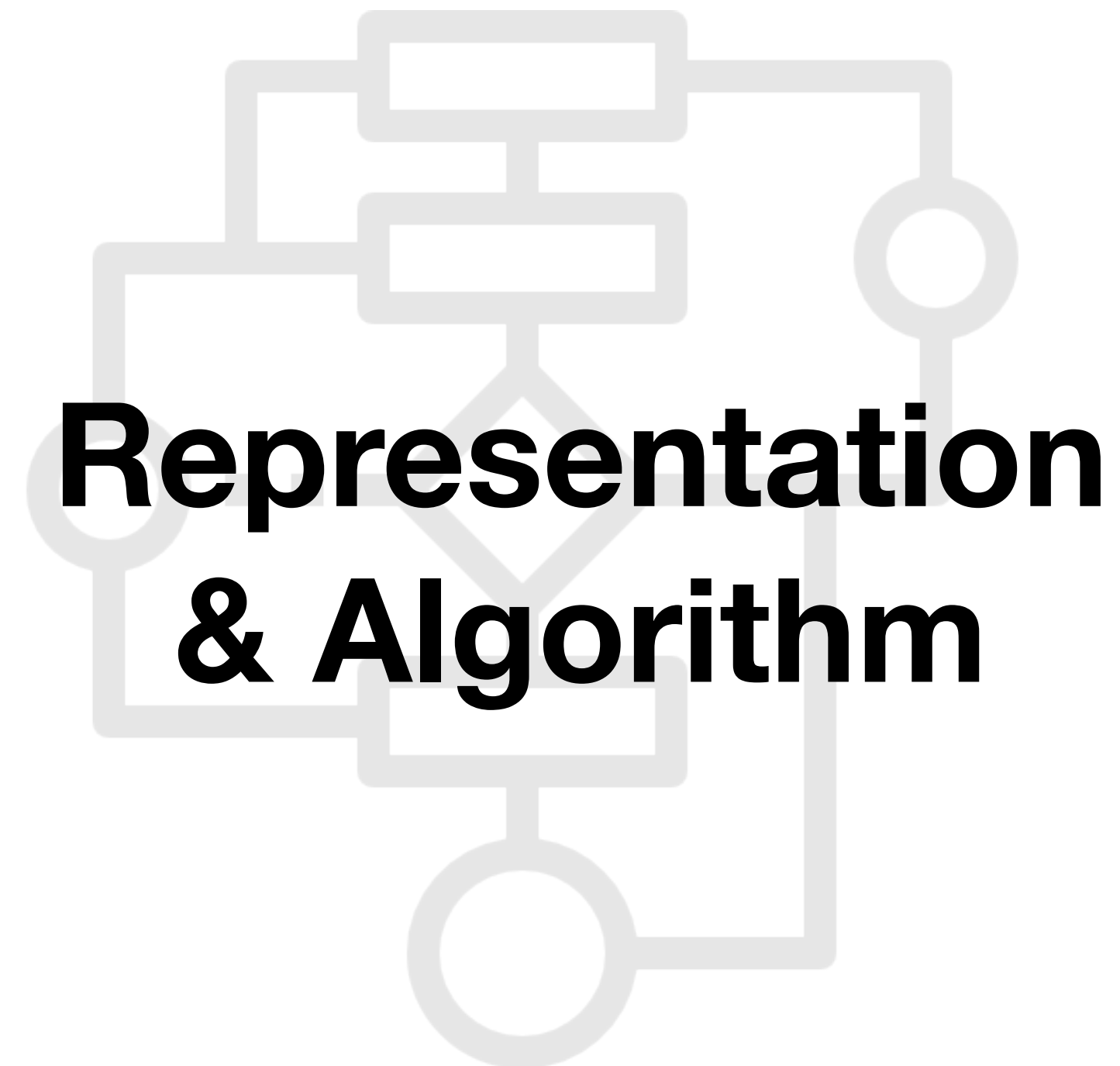
Selection in  
human culture



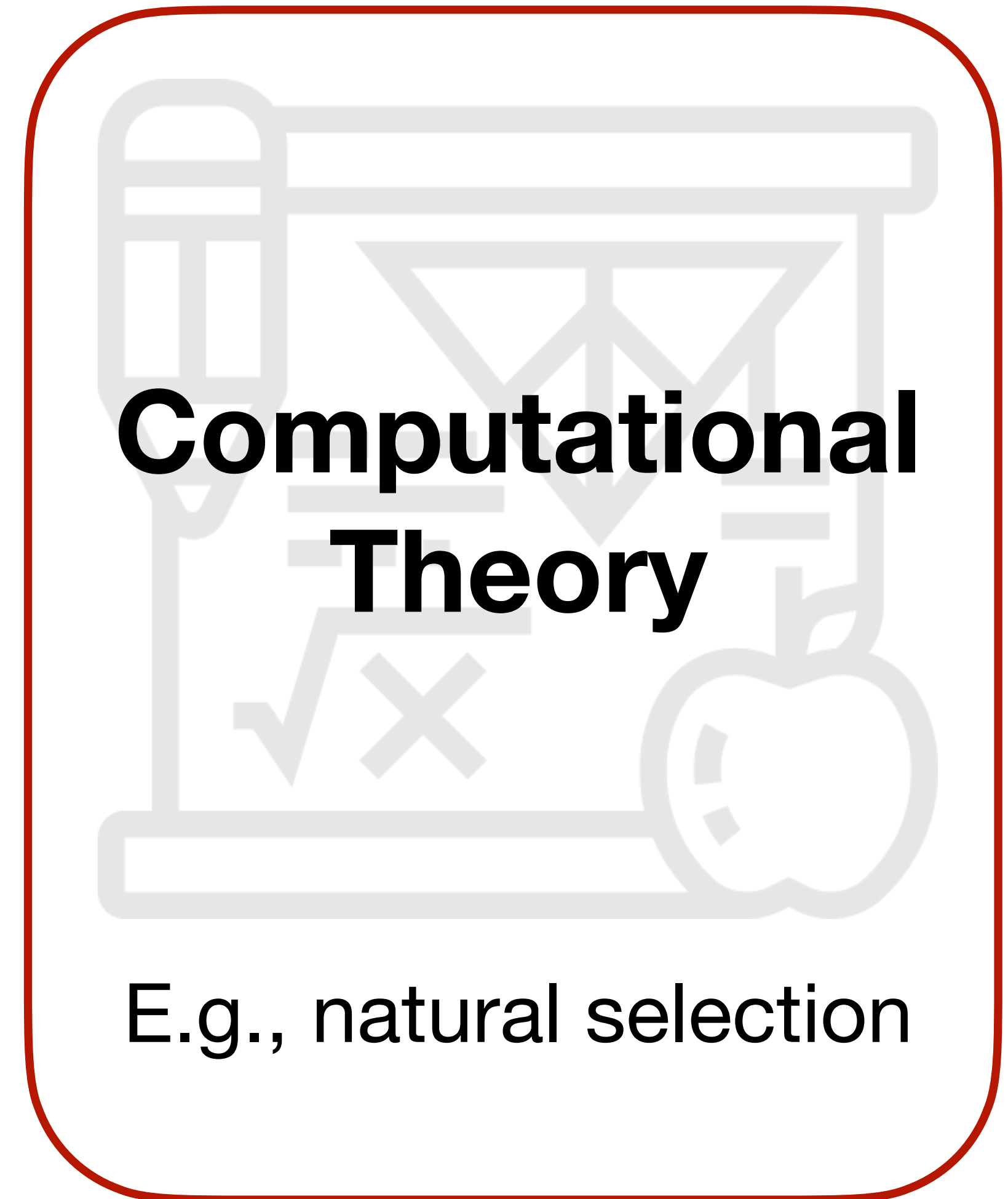
# Evolution as a Computational Principle



E.g., gene and fossils



E.g., mechanism for  
sexual dimorphism, meme



E.g., natural selection



# Common Misunderstanding

**Q:** Evolution as an optimization (i.e., optimizing the fitness)!?

**A:** Not really!

*“Evolution has no long-term goal.*

*There is no long-distance target,*

*no final perfection to serve as a criterion for selection.”*

*– Richard Dawkins*



**Lecture III.c**  
(Jan. 20 10am-10:50am ET)



# Neuroscience

*“There is no scientific study more vital to man than the study of his own brain. Our entire view of the universe depends on it.”*

*– Francis Crick*

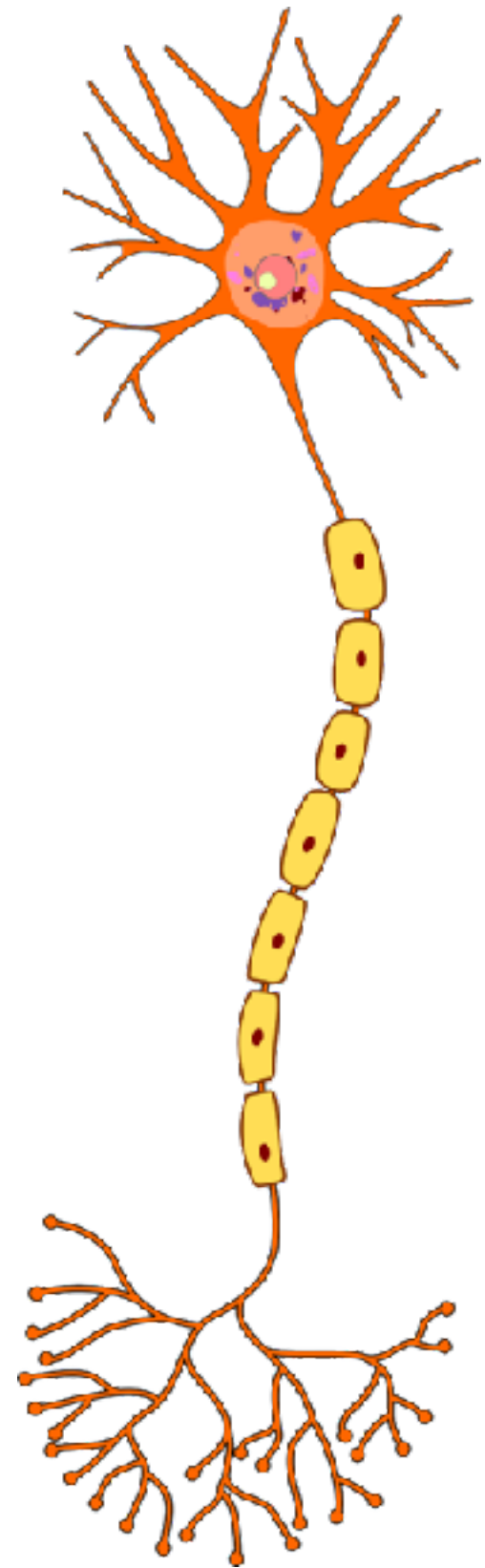


# The Biggest Mystery

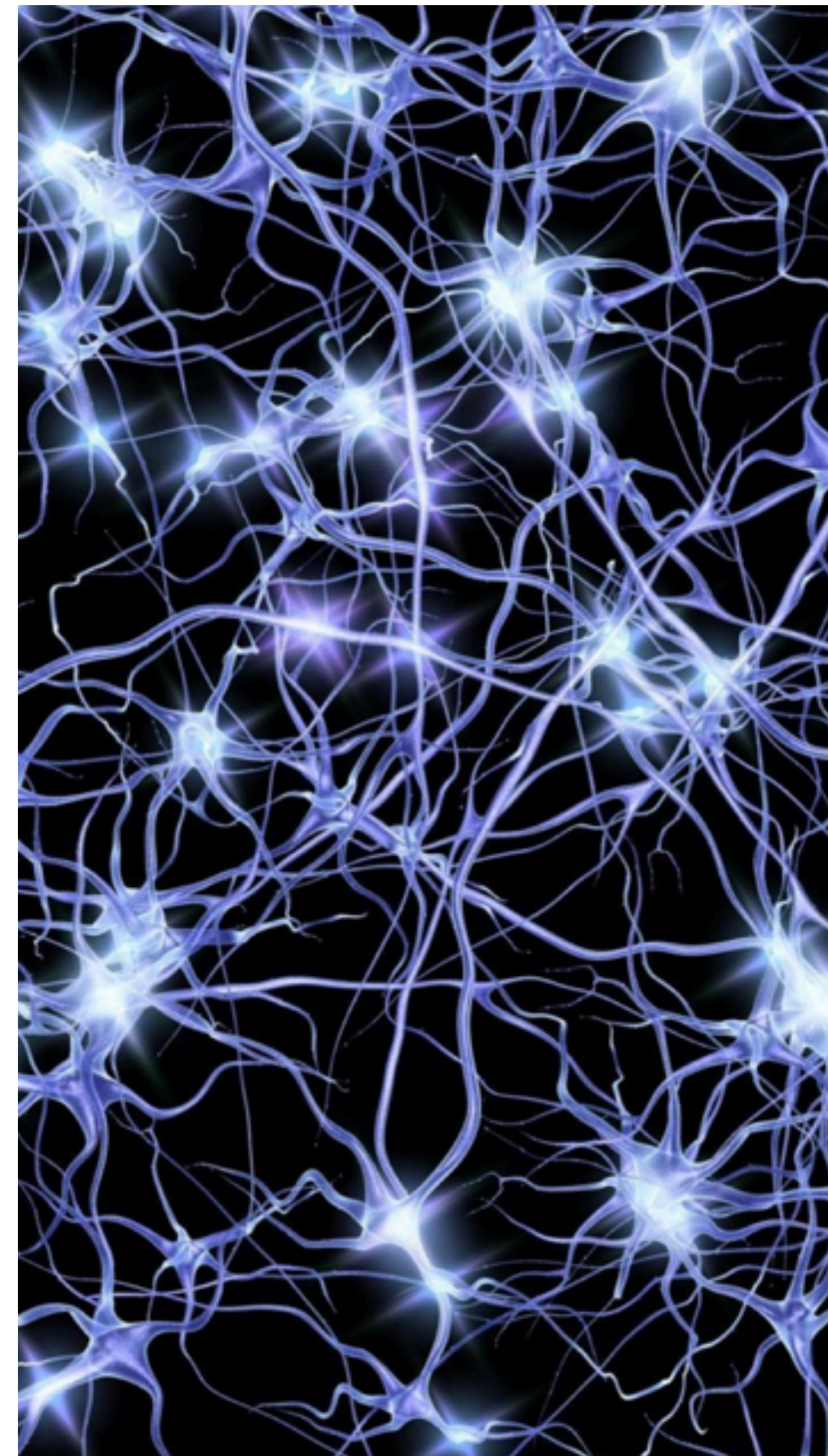




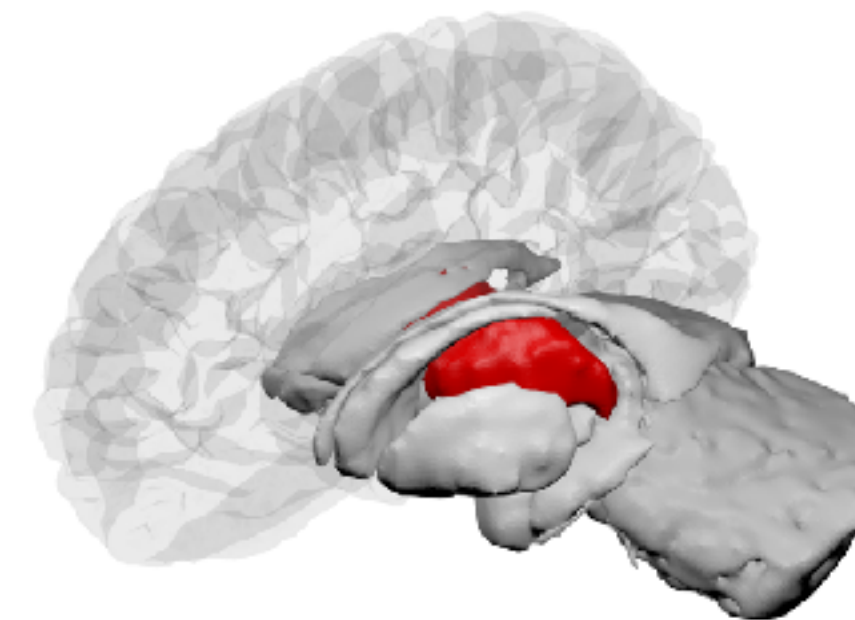
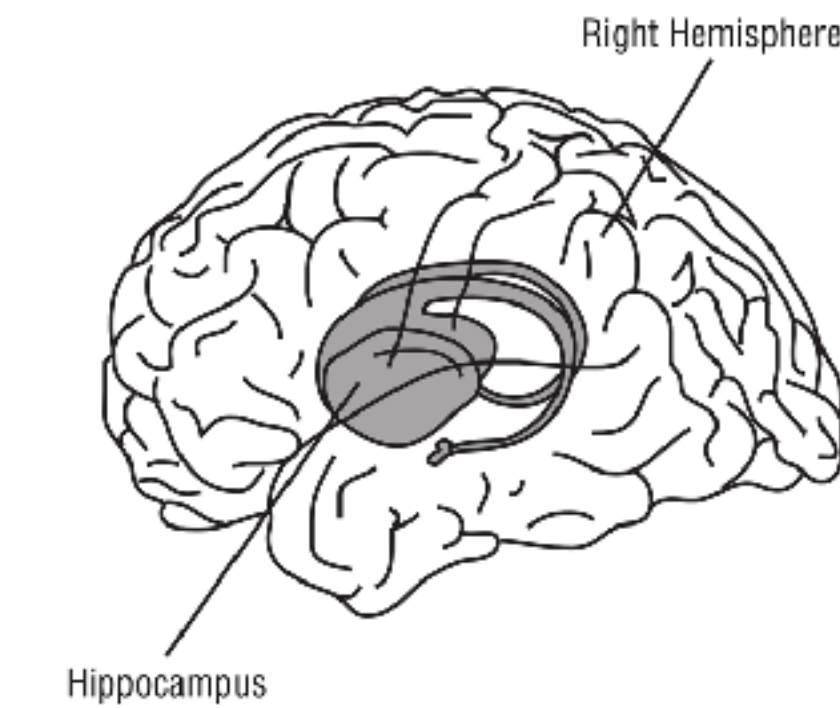
# From Neuron to Consciousness



# Single neuron



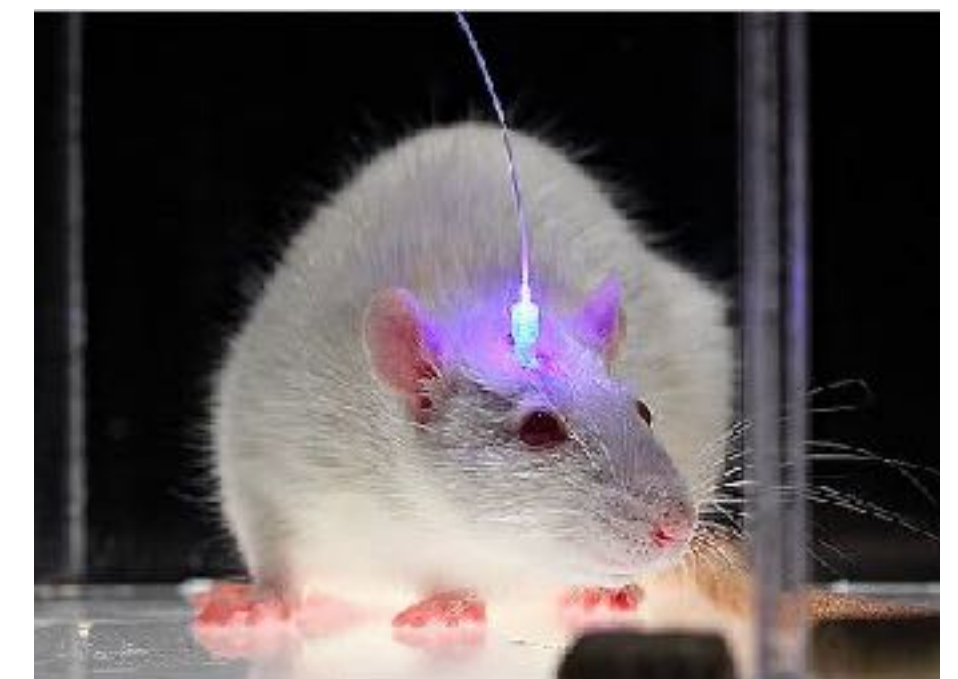
# Network



# System



# Consciousness



# Behavior



# Many Different Approaches in Neuroscience

*“Our researchers often cross the boundaries of established fields, or invent new disciplines entirely. Conceptually, however, we think of our research in four broad categories:”*

*– MIT Department of Brain and Cognitive Sciences*

> Cellular and Molecular Neuroscience



> Systems Neuroscience



> Cognitive Science

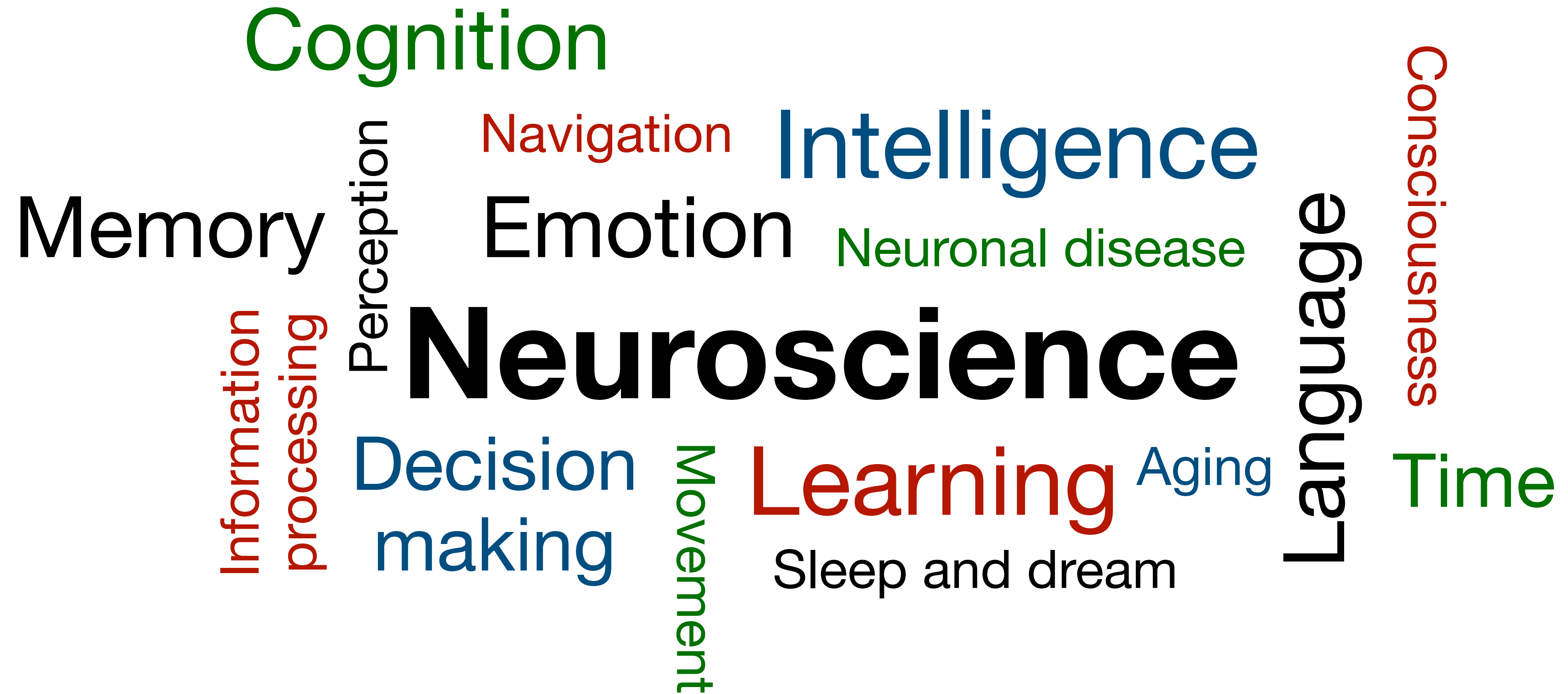


> Computational Neuroscience





# Computations in the Brain?

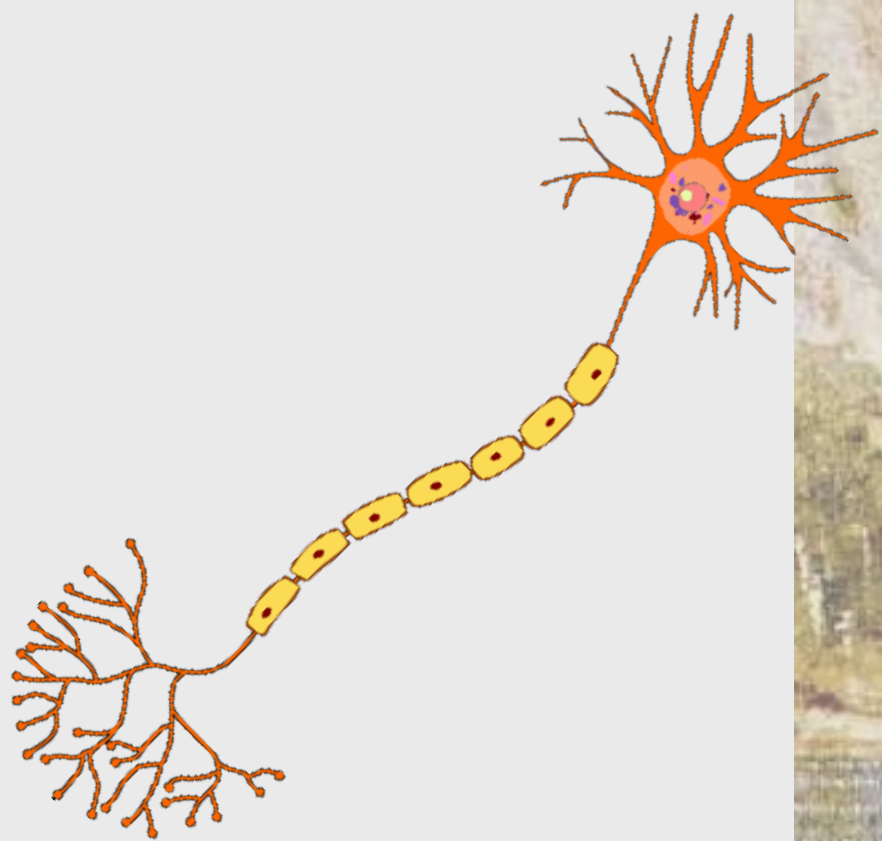


So many aspects mingle together, how do we start!?

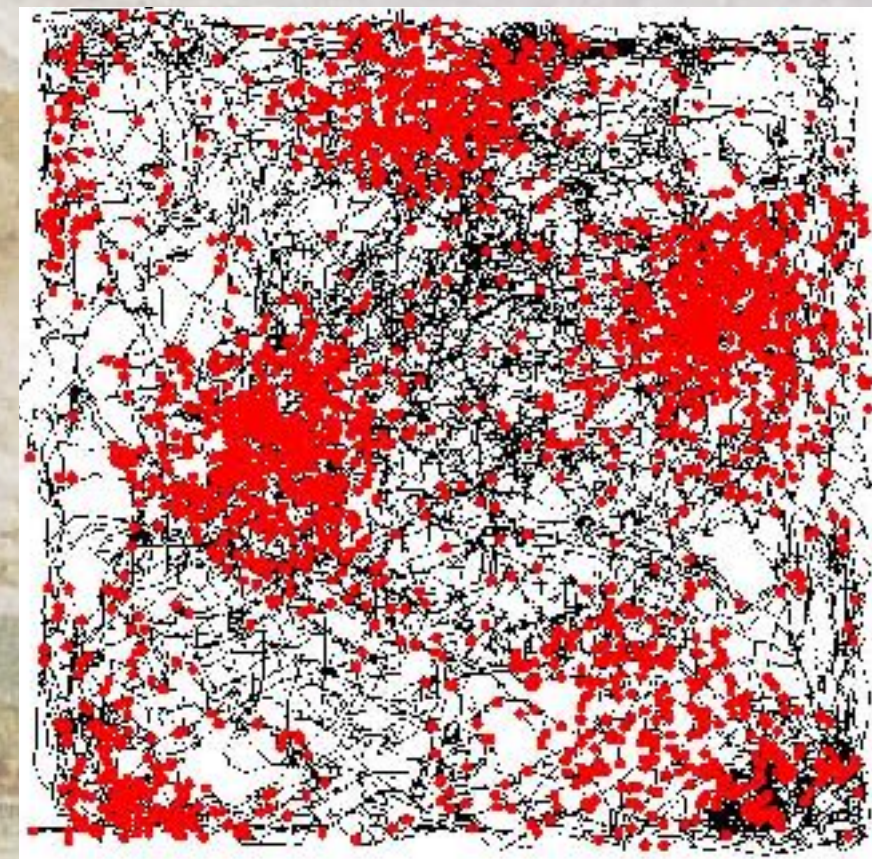


# Beagle Expedition in Neuroscience!?

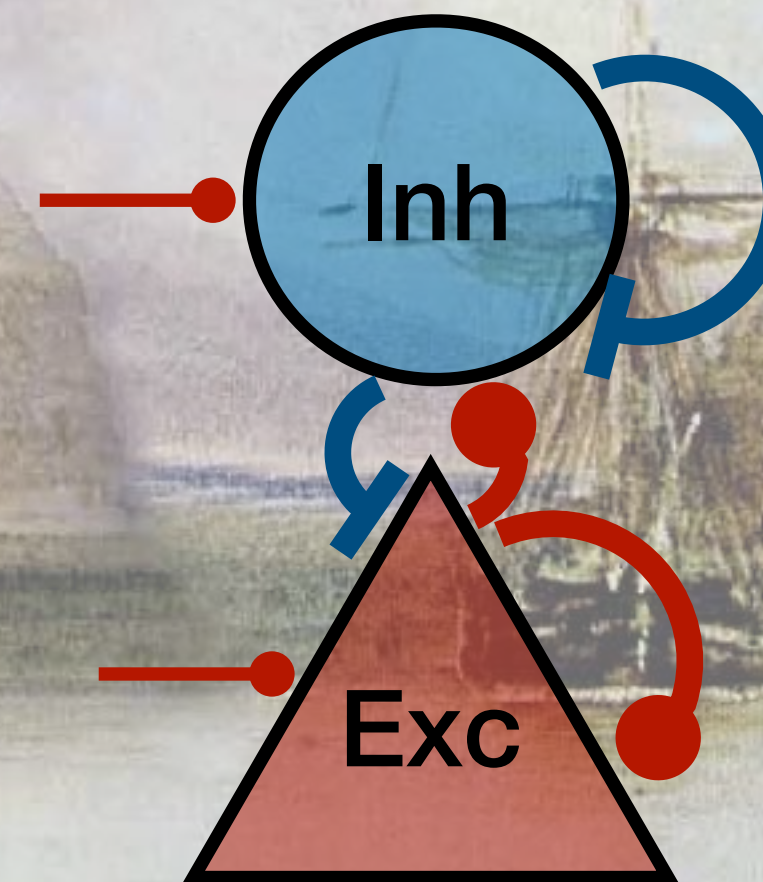
## Example 1: Single Neuron



## Example 2: Grid Cells



## Example 3: Inhibition Stabilized Networks



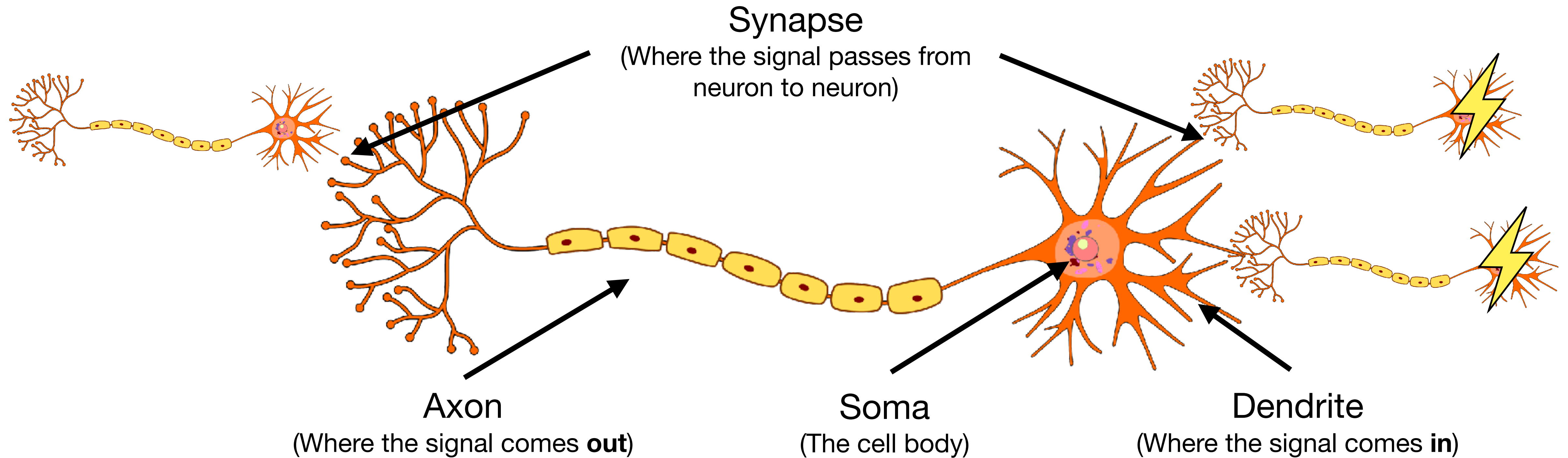
## More!



Brabeeba Wang  
(Jan. 20 11am-12pm ET)



# Example 1: Single Neuron



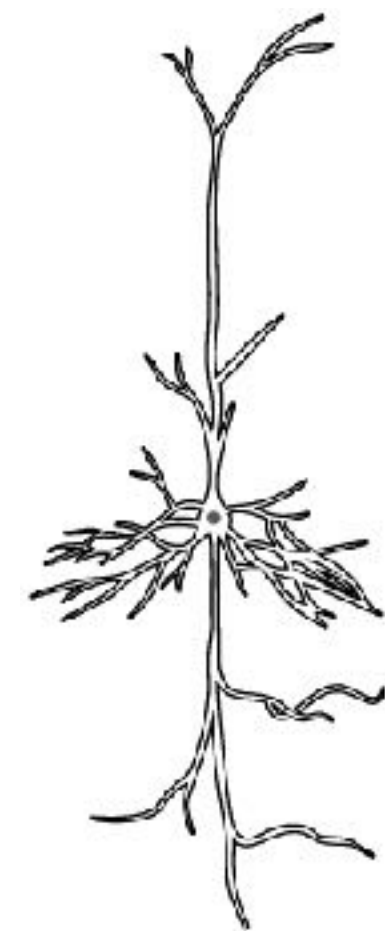
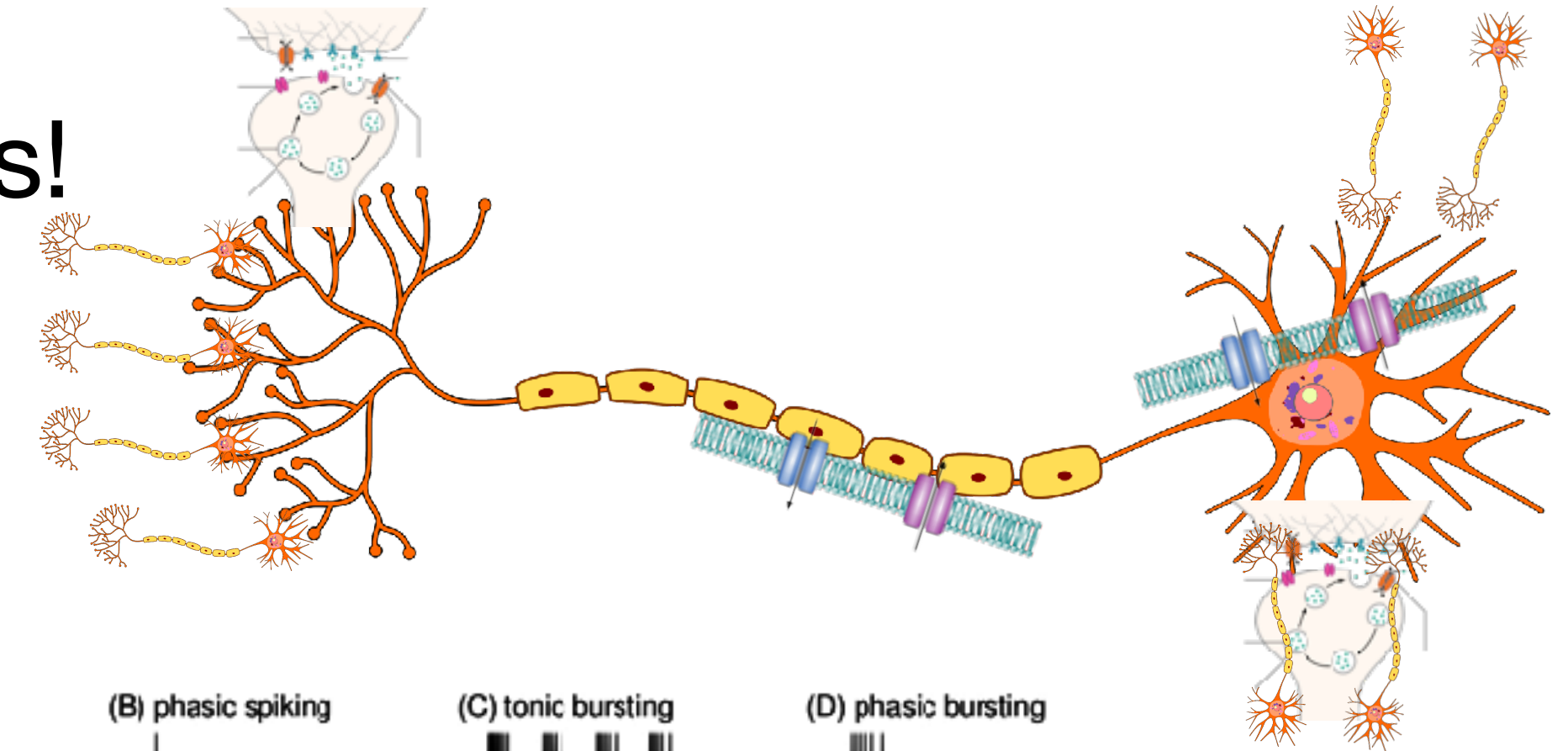
This is oversimplified, in real brain there are special cases and are ubiquitous!



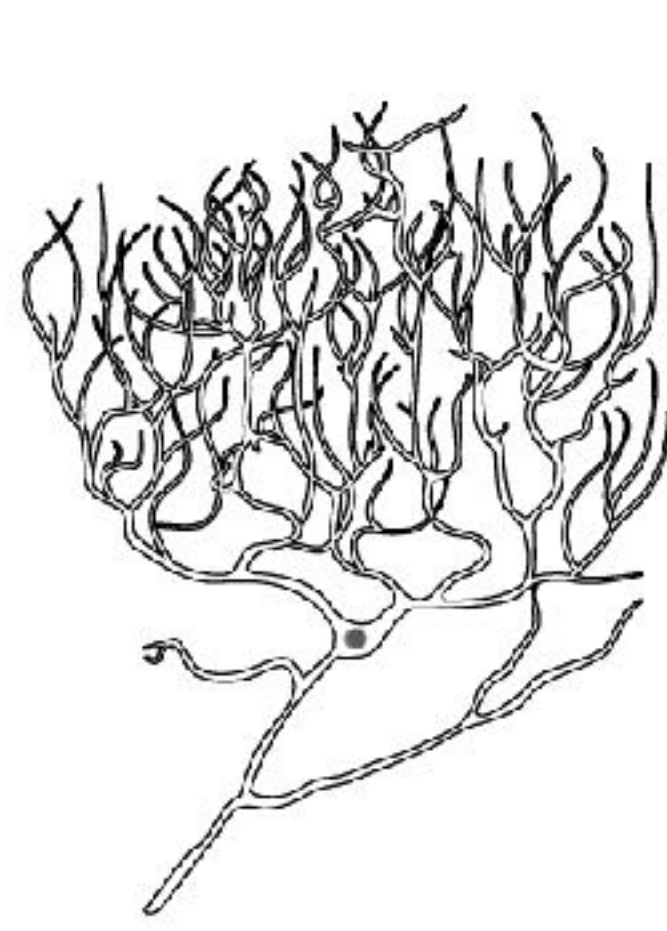
# Neurons Can be Complicated and Diverse

There are many factors decide neuron's activities!

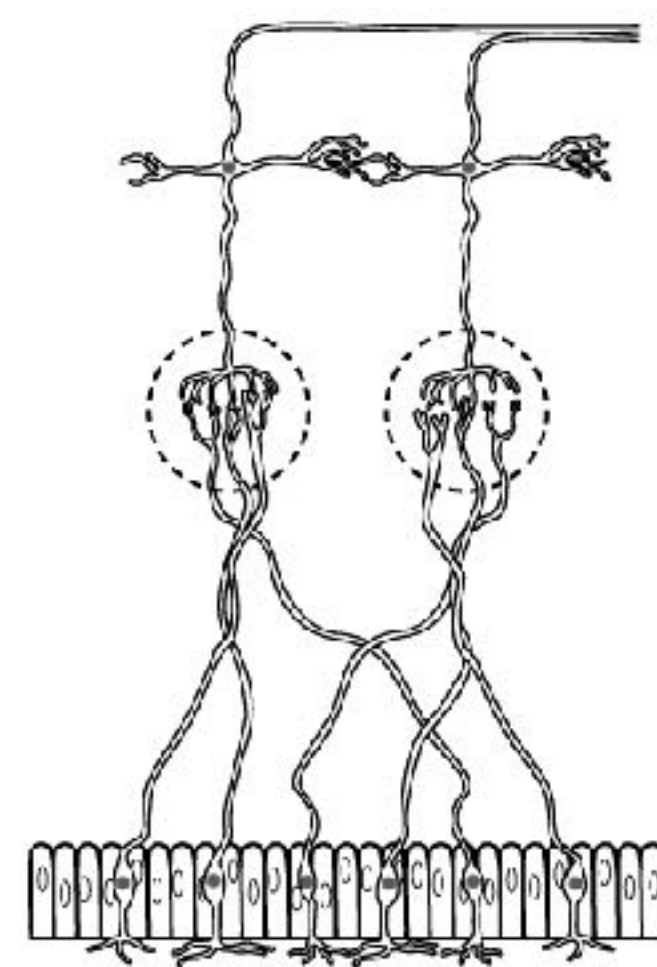
- Ion channel (interaction with the outside environment).
- Neurotransmitter (interaction with other neurons).
- Axon and dendrite (geometry of the interconnection).
- ...



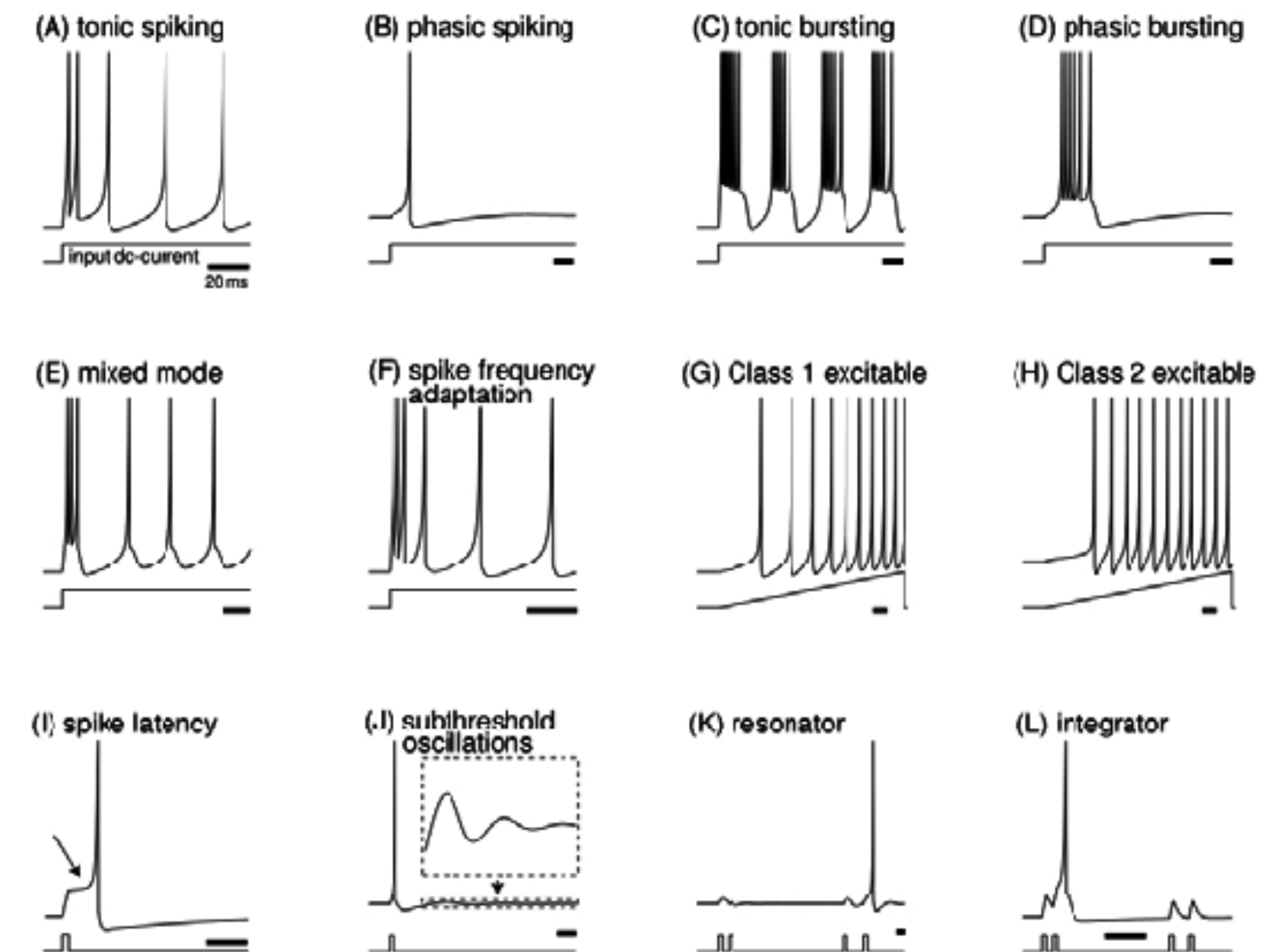
(a) Pyramidal cell of the cerebral cortex



(b) Purkinje cell of the cerebellar cortex



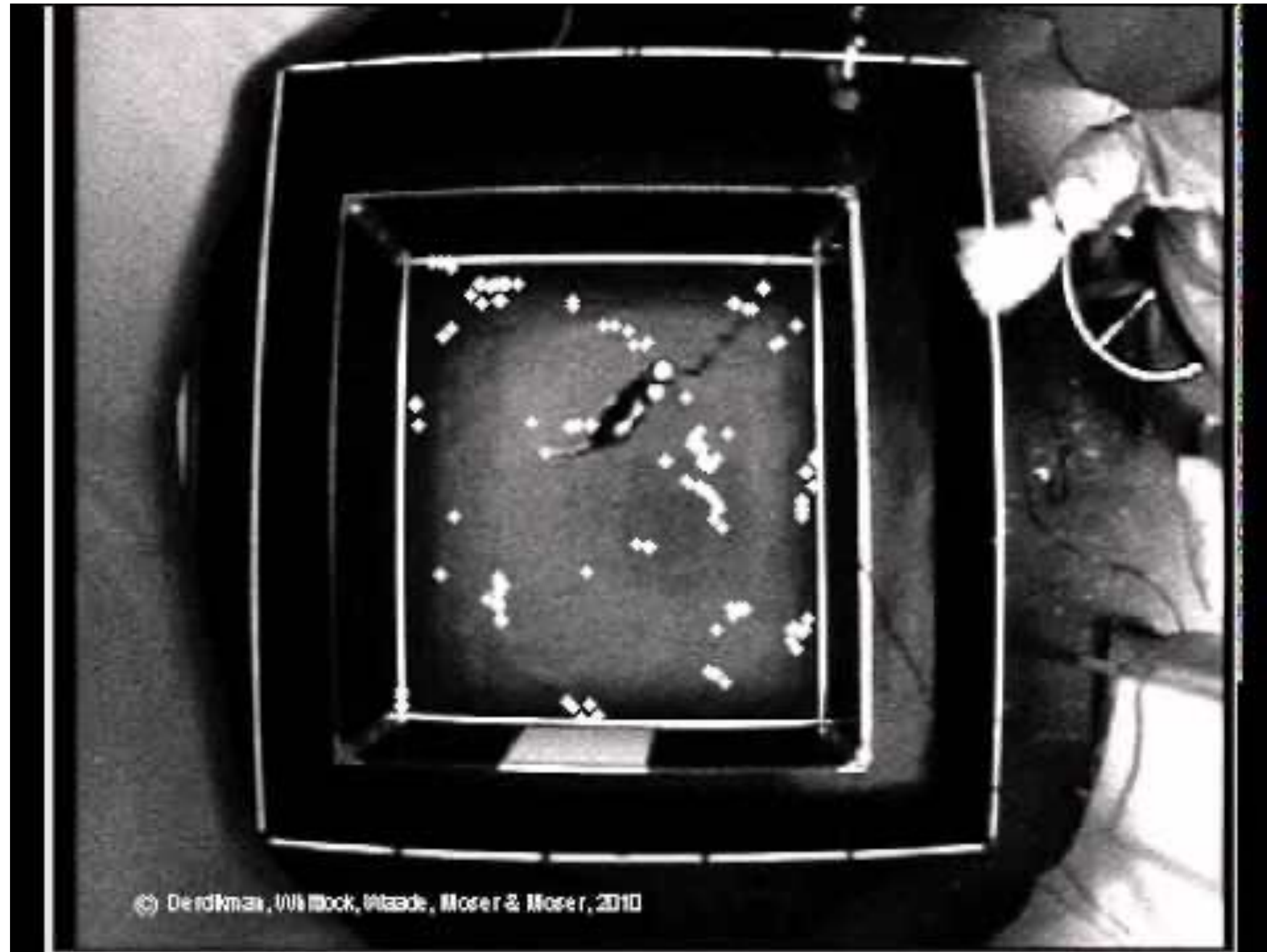
(c) Olfactory cells in the olfactory epithelium and olfactory bulbs



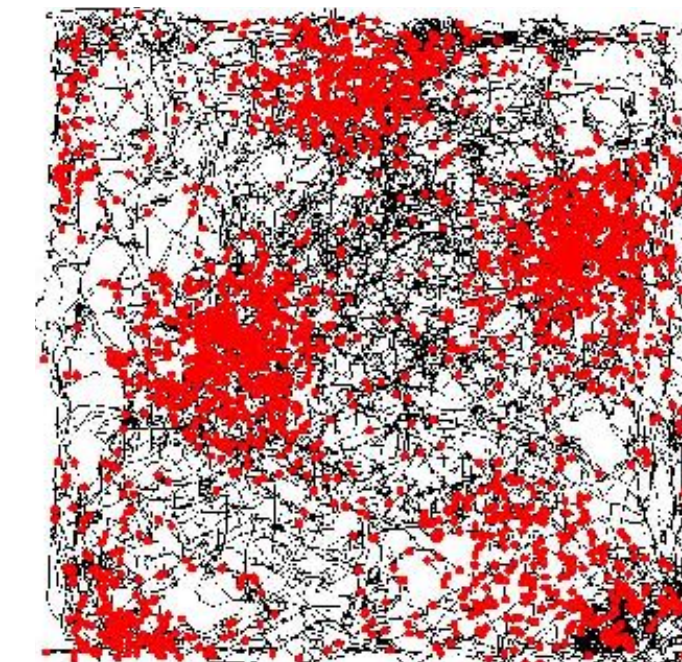
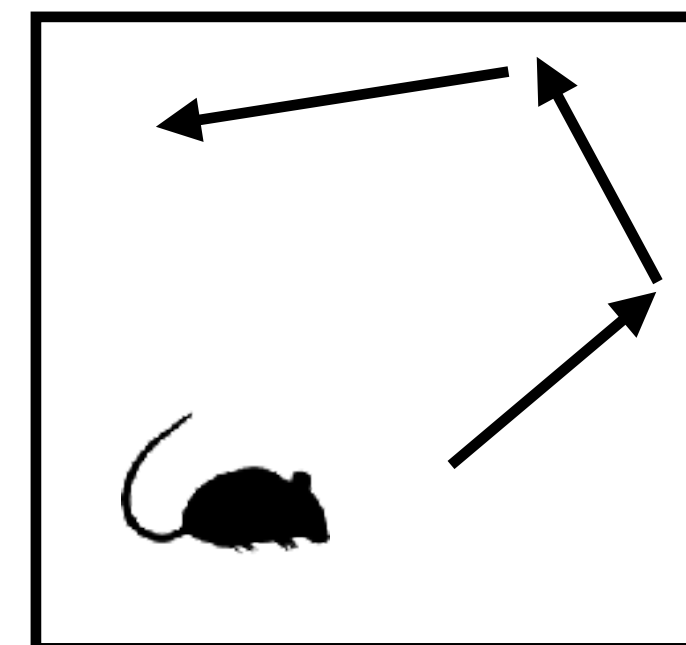
Single neuron seems to do lots of computation already!?



# Example 2: Grid Cells



The 2014 Nobel Prize in Physiology or Medicine

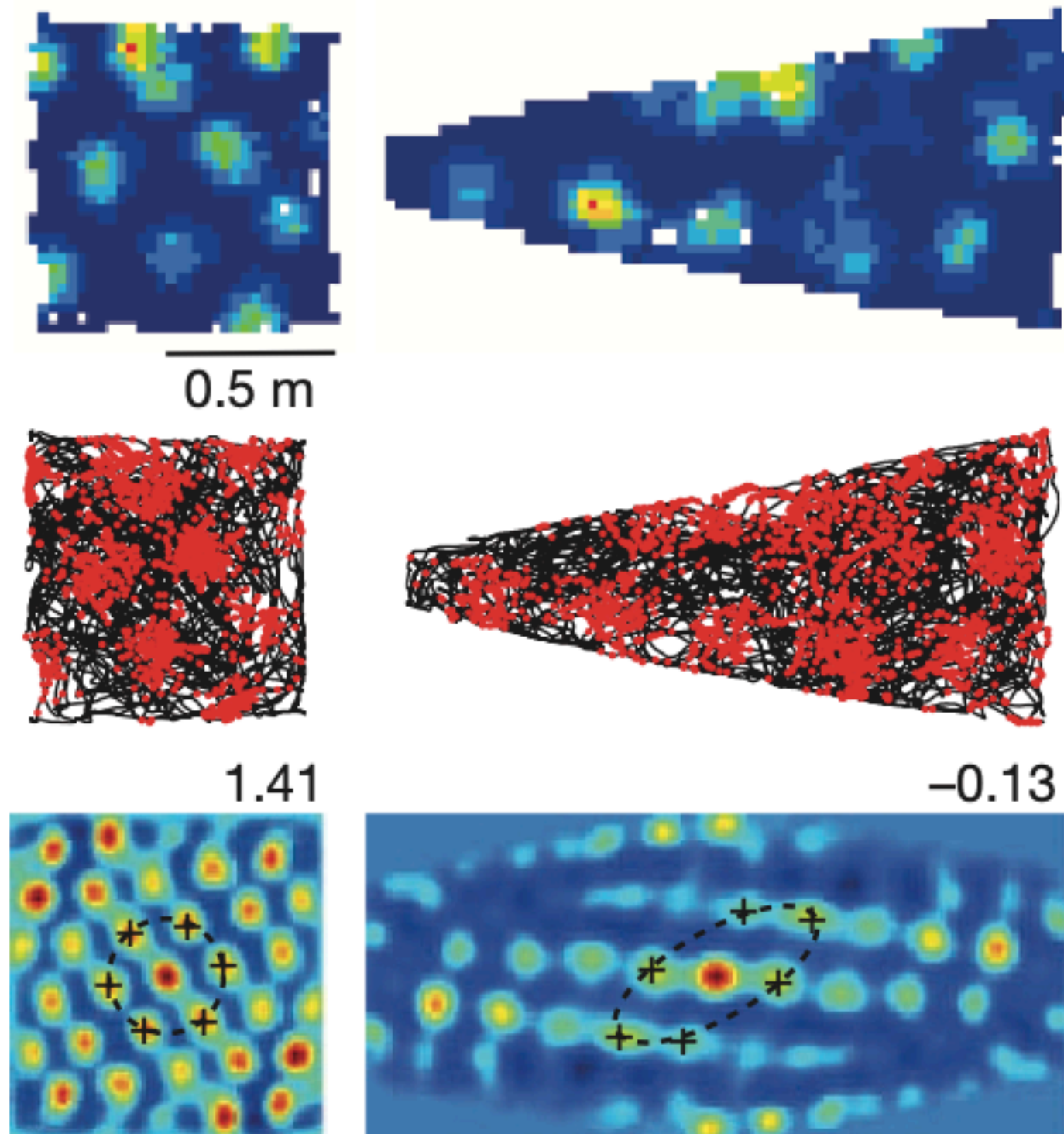


Receptive field of a single grid cell

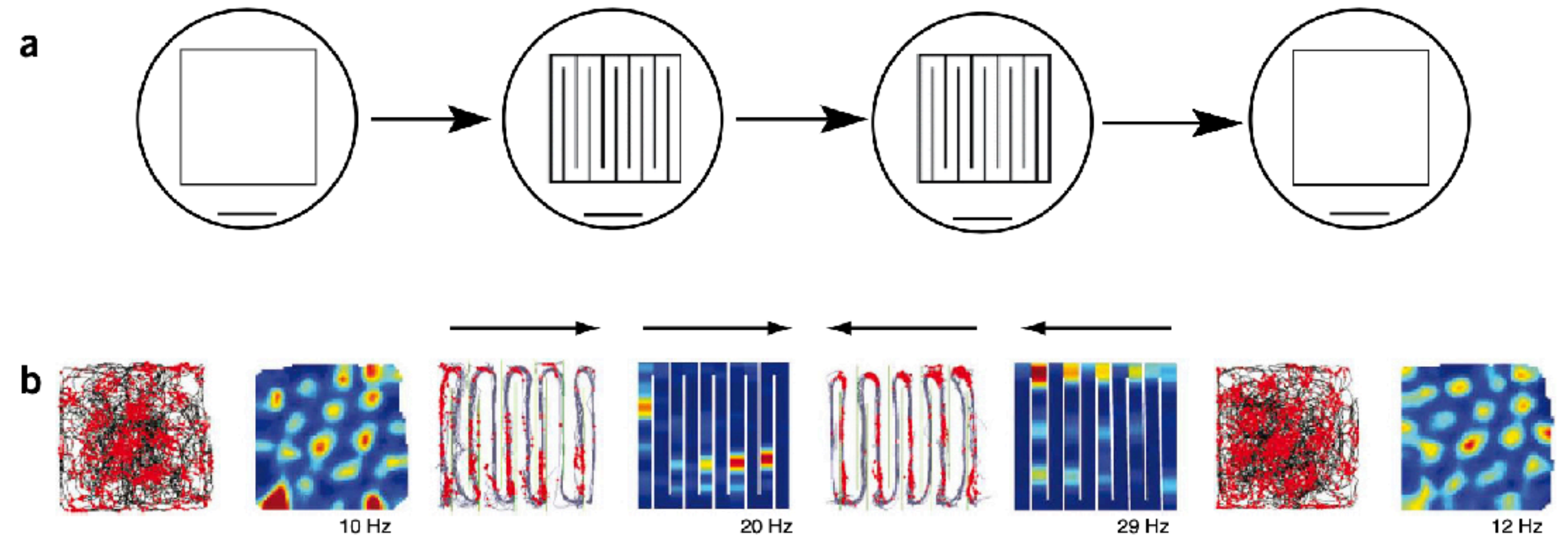
This is oversimplified, in real mice there are many special cases!



# Not Always Hexagonal Grids!



\* Krupic, Julija, et al. "Grid cell symmetry is shaped by environmental geometry." *Nature* 518.7538 (2015): 232-235.



\* Derdikman, Dori, et al. "Fragmentation of grid cell maps in a multicompartiment environment." *Nature neuroscience* 12.10 (2009): 1325-1332.

**Q:** Why and how grid cells emerge?

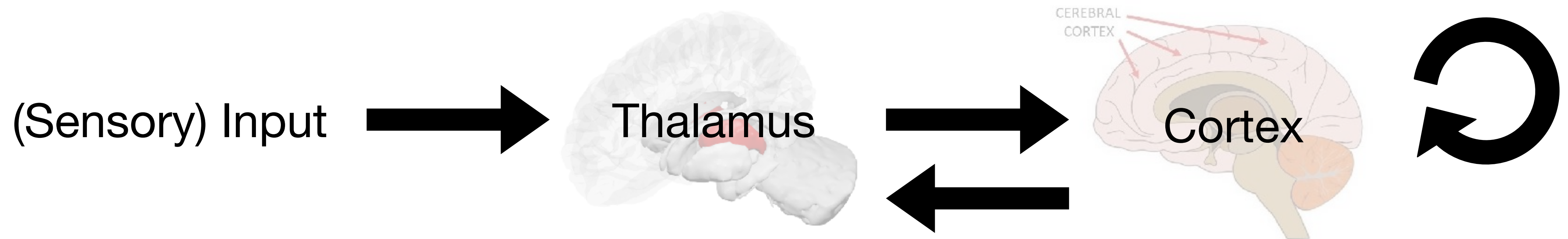
**Q:** What computation are grid cells doing?

**Q:** What's the role of grid cells in the brain?



# Example 3: Inhibition Stabilized Networks

*Toward understanding cortical computation*

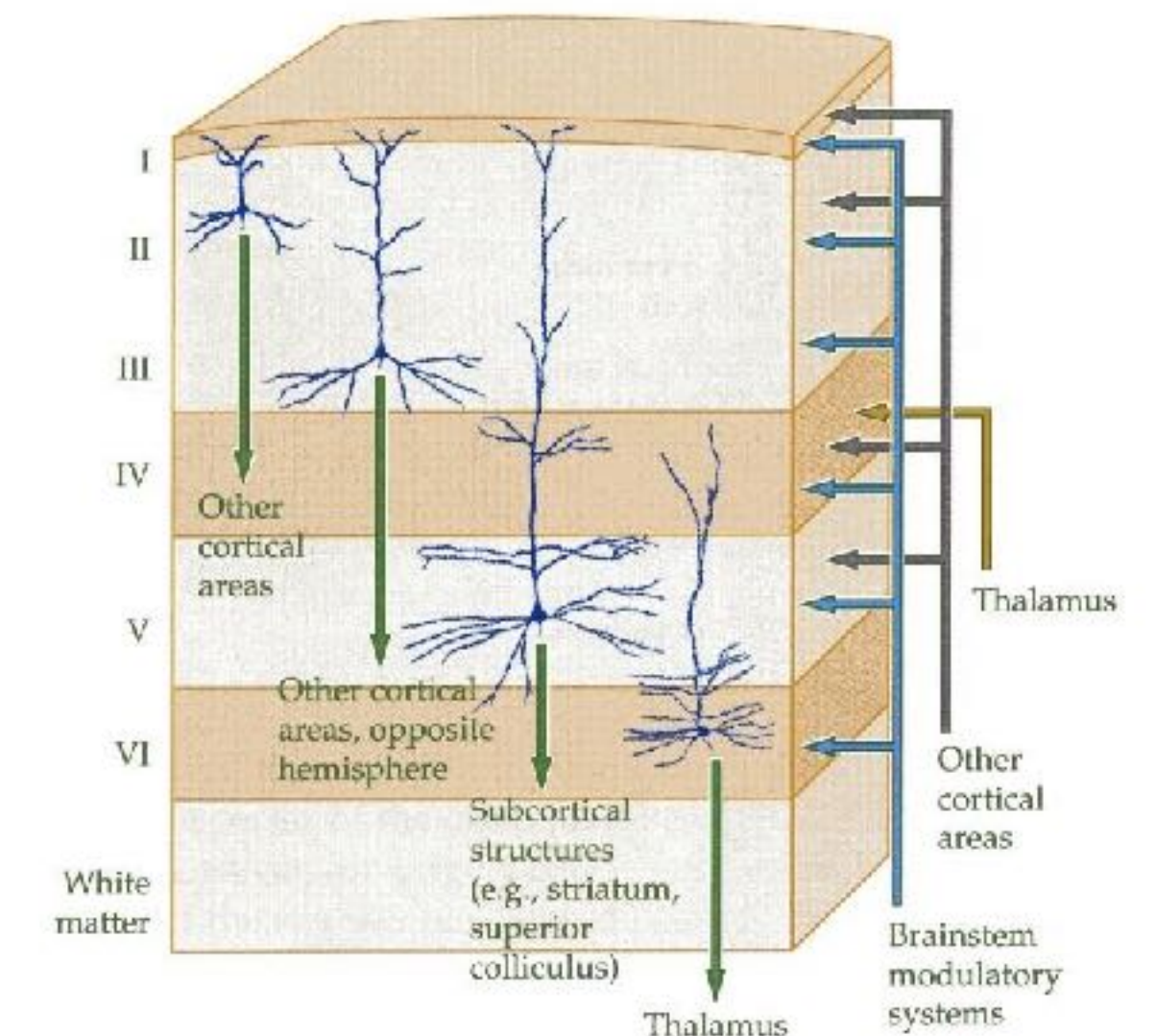


Cortex plays a key role in lots of aspects of computation!

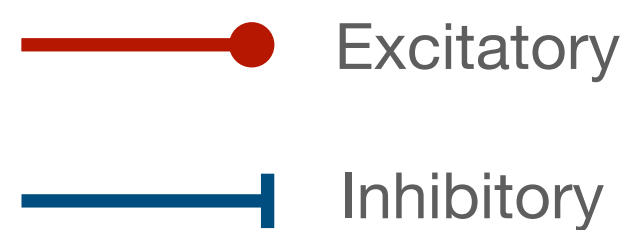
**Q:** How does cortex reconcile the feedforward and recurrent interactions?

Meanwhile, cortex is very structured!

**Q:** Network motifs and cortical sub-functions?

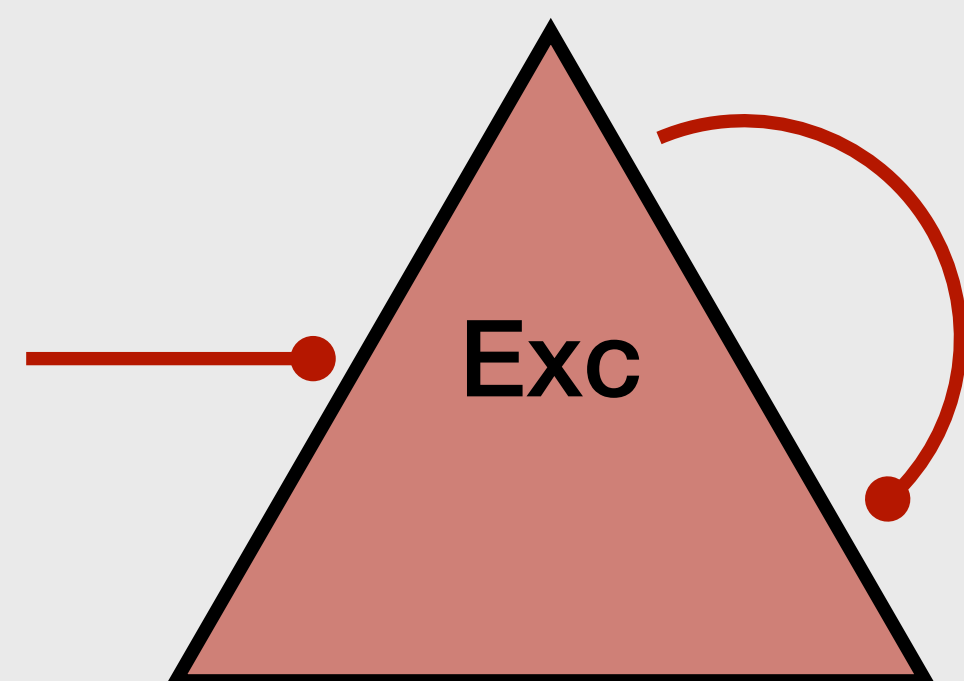






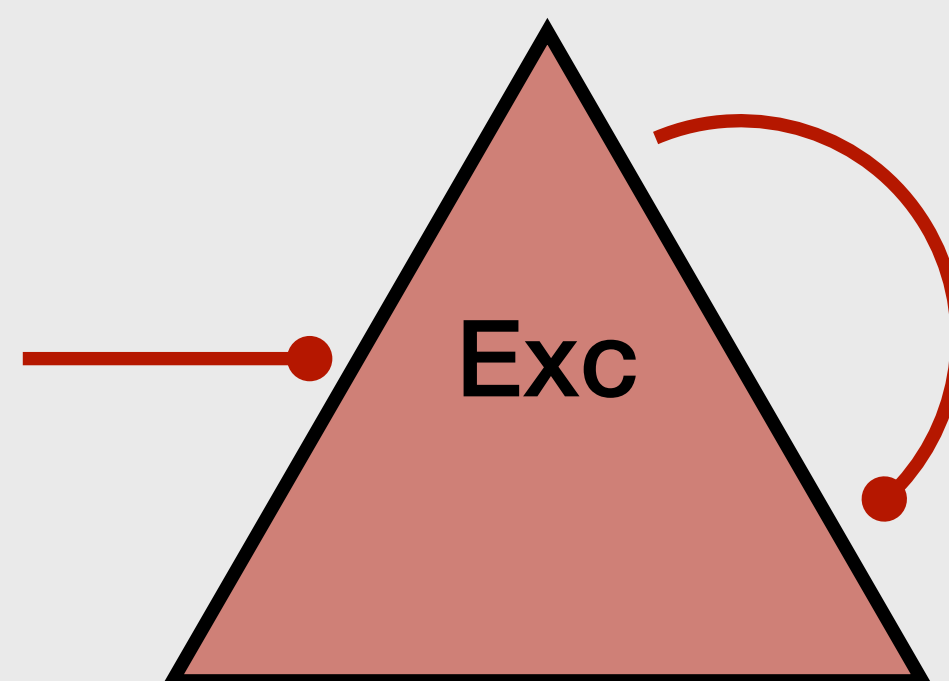
# Network Motifs Proposed by Theory

## Selecting Patterns



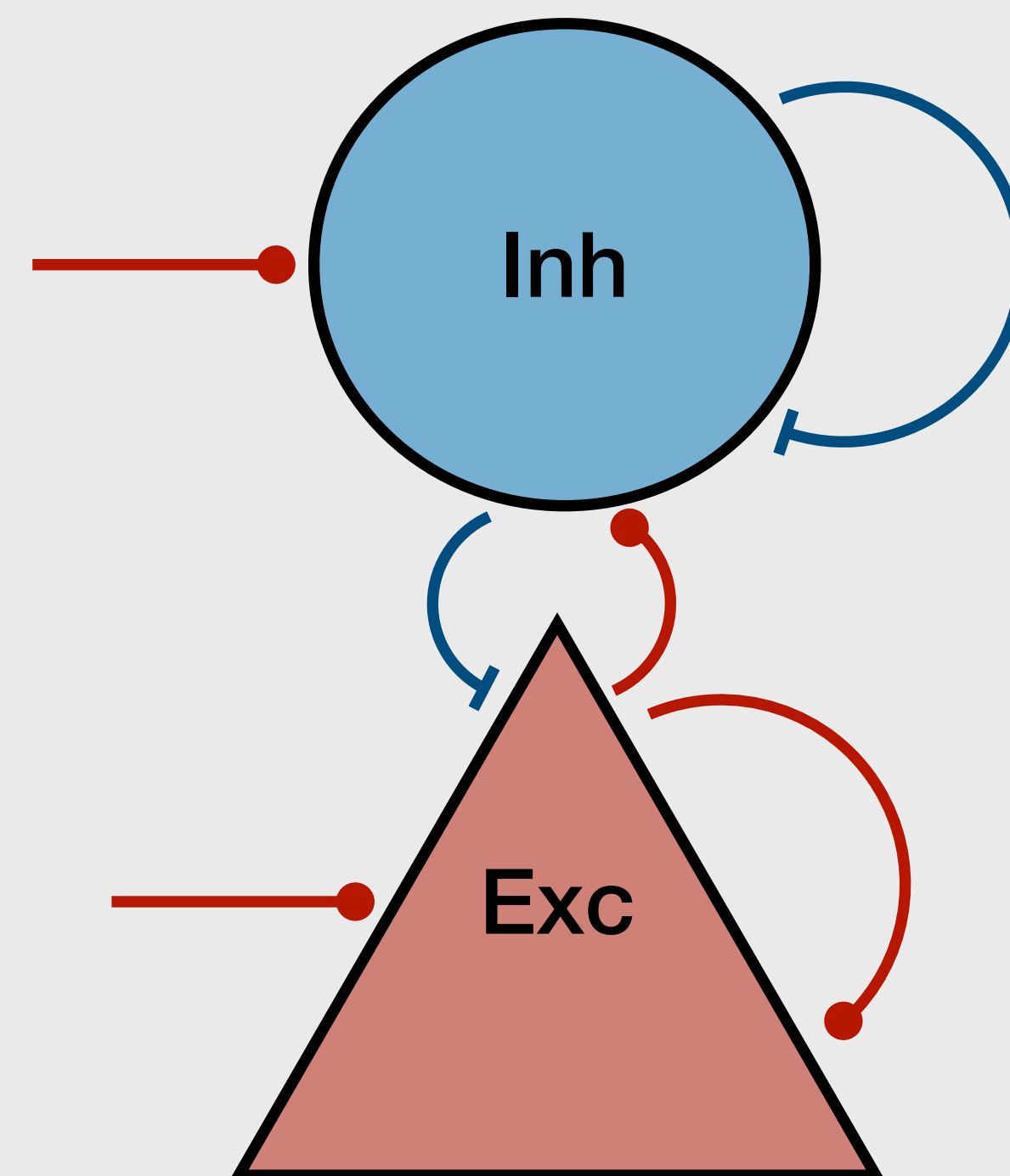
Strong excitatory recurrence constraints the set of stable activity patterns.

## Amplifying Inputs



Strong excitatory recurrence amplifies the feedforward inputs (e.g., from thalamus).

## E/I Balance Network

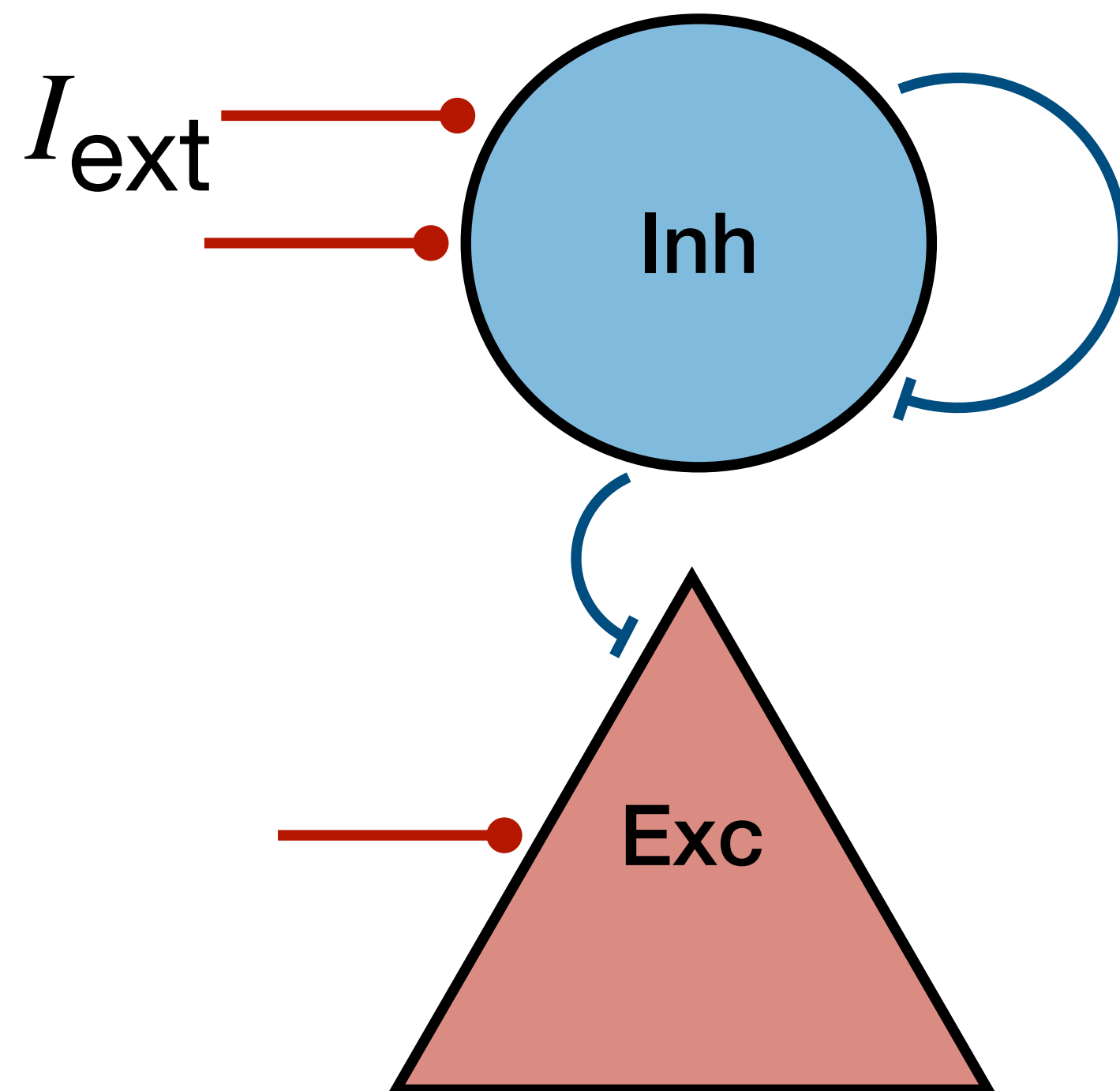


Variability of cortical spiking.

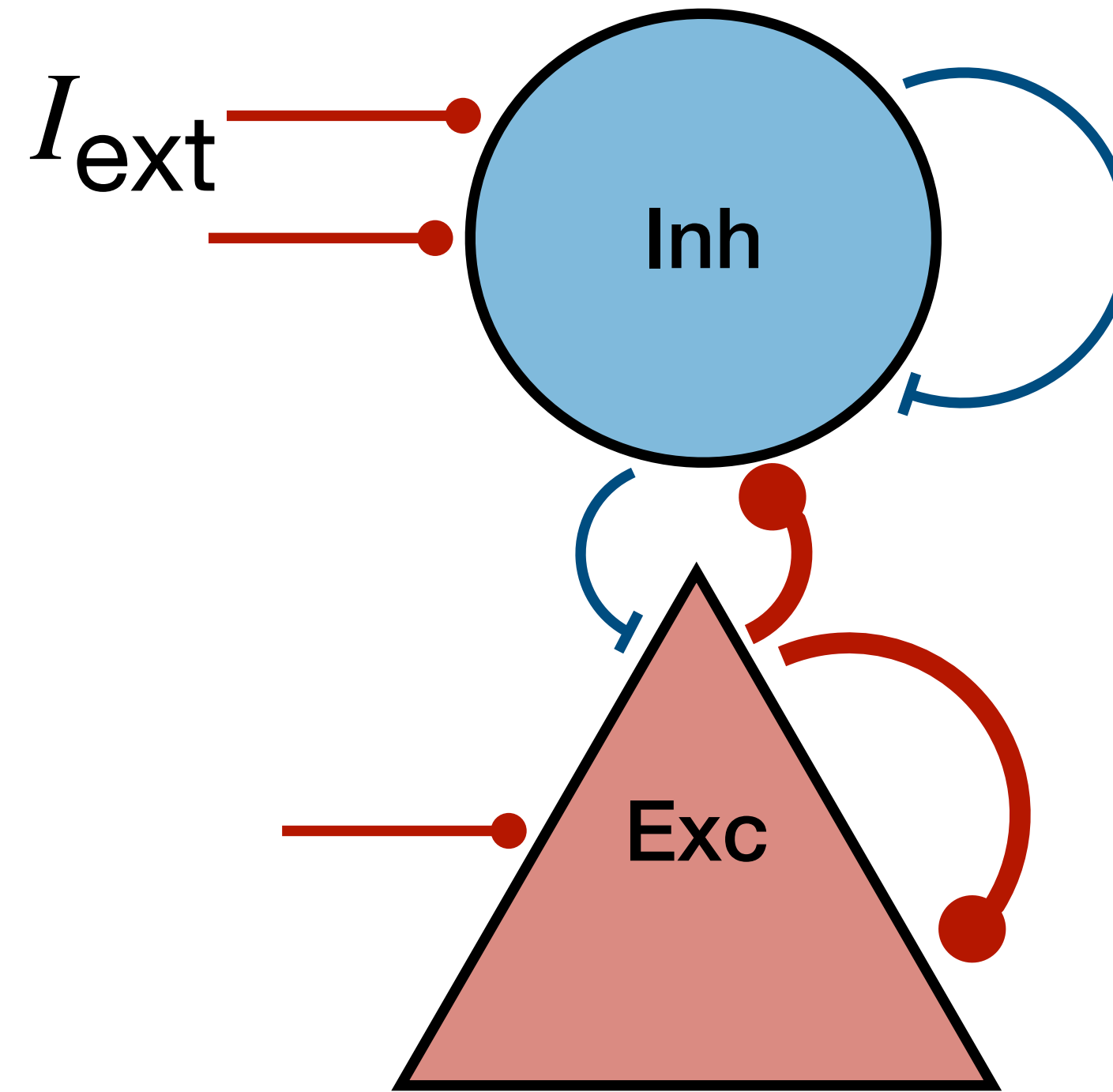


# Connectivity Strengths Matter!

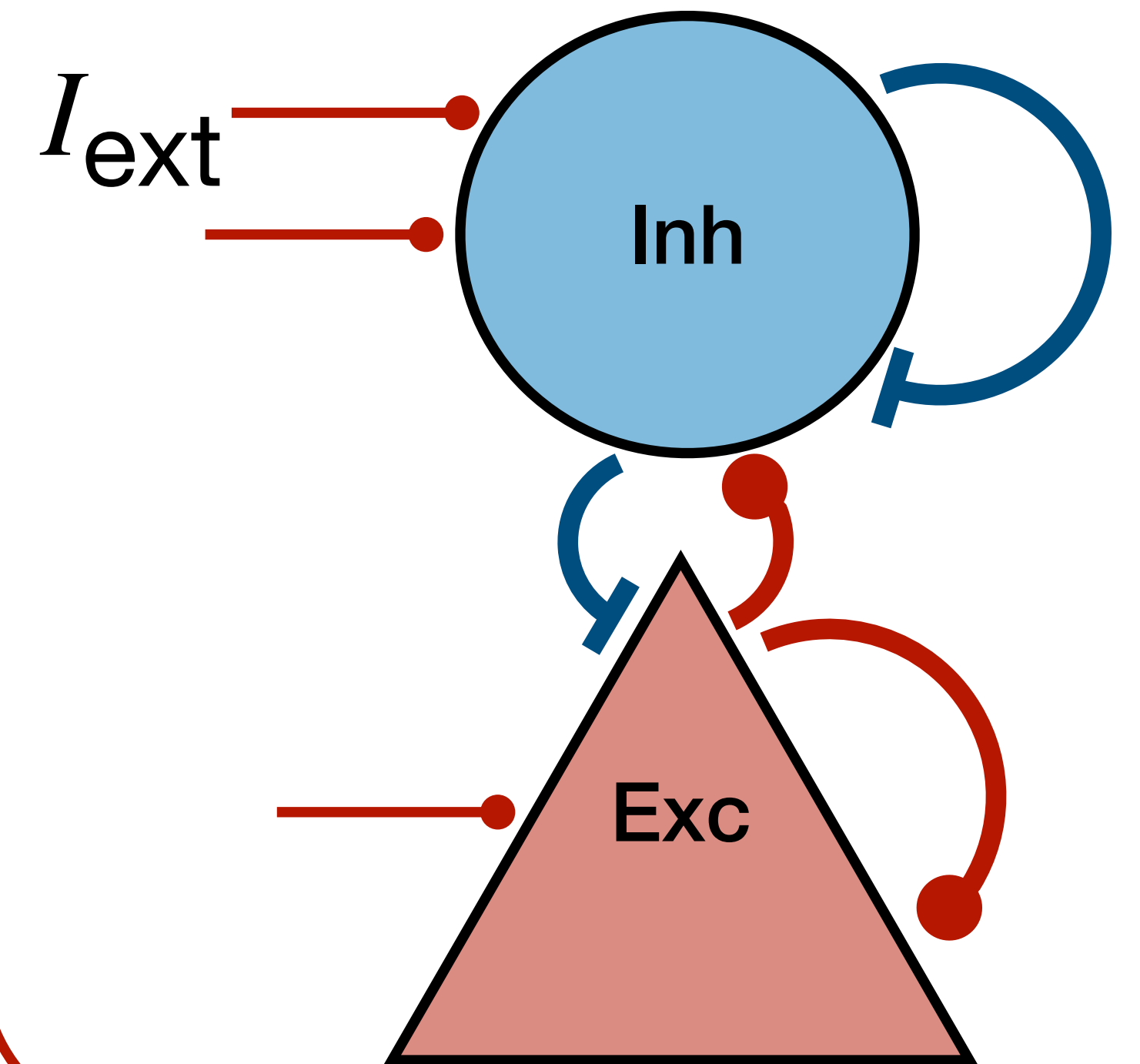
Feedforward Inhibition



Recurrent Inhibition



Inhibition-Stabilized



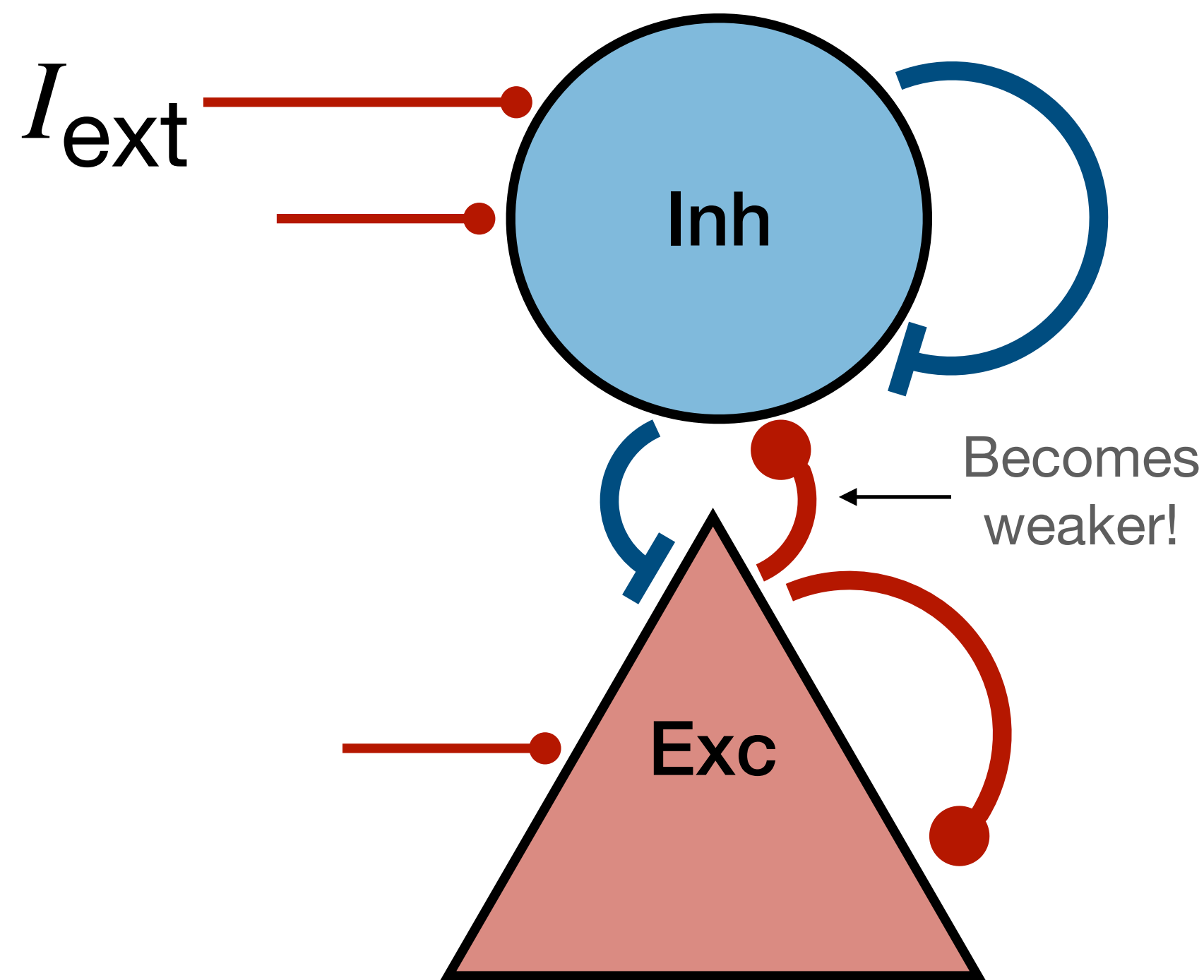
Different connectivity strengths induces different activity patterns!



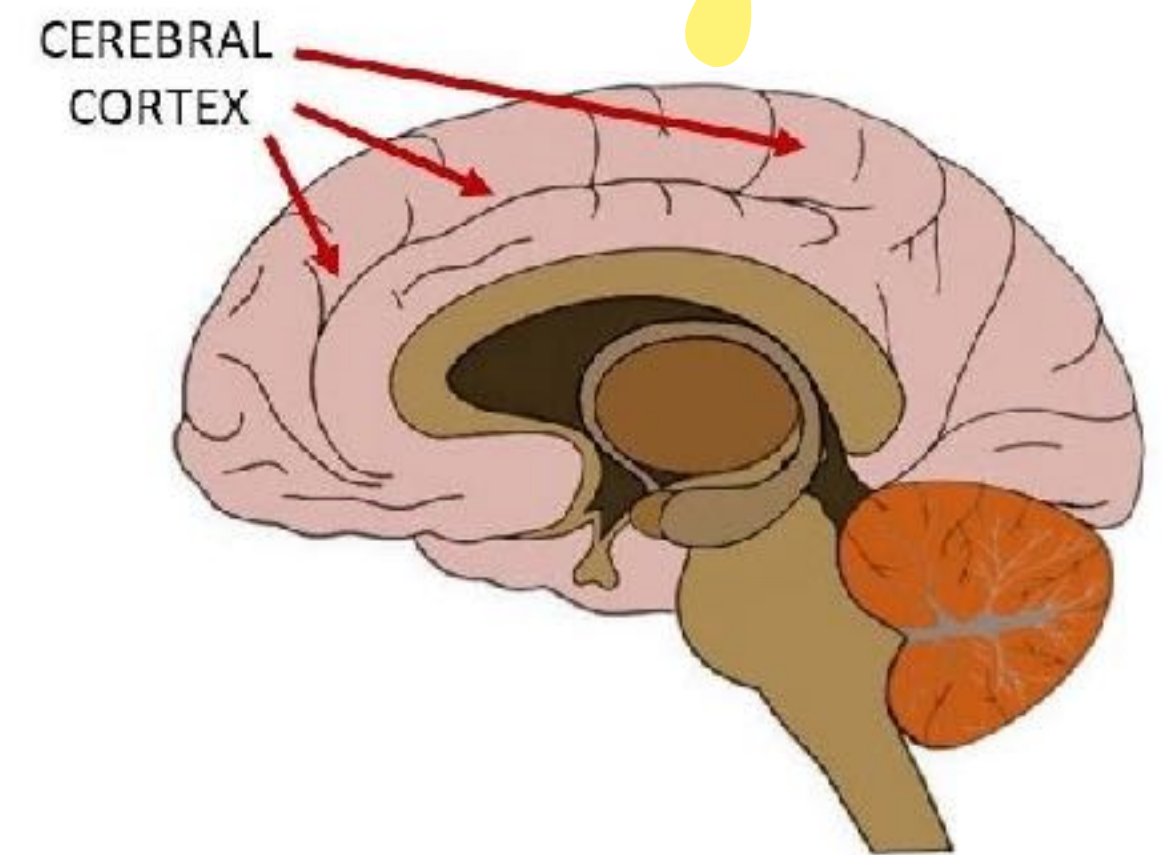
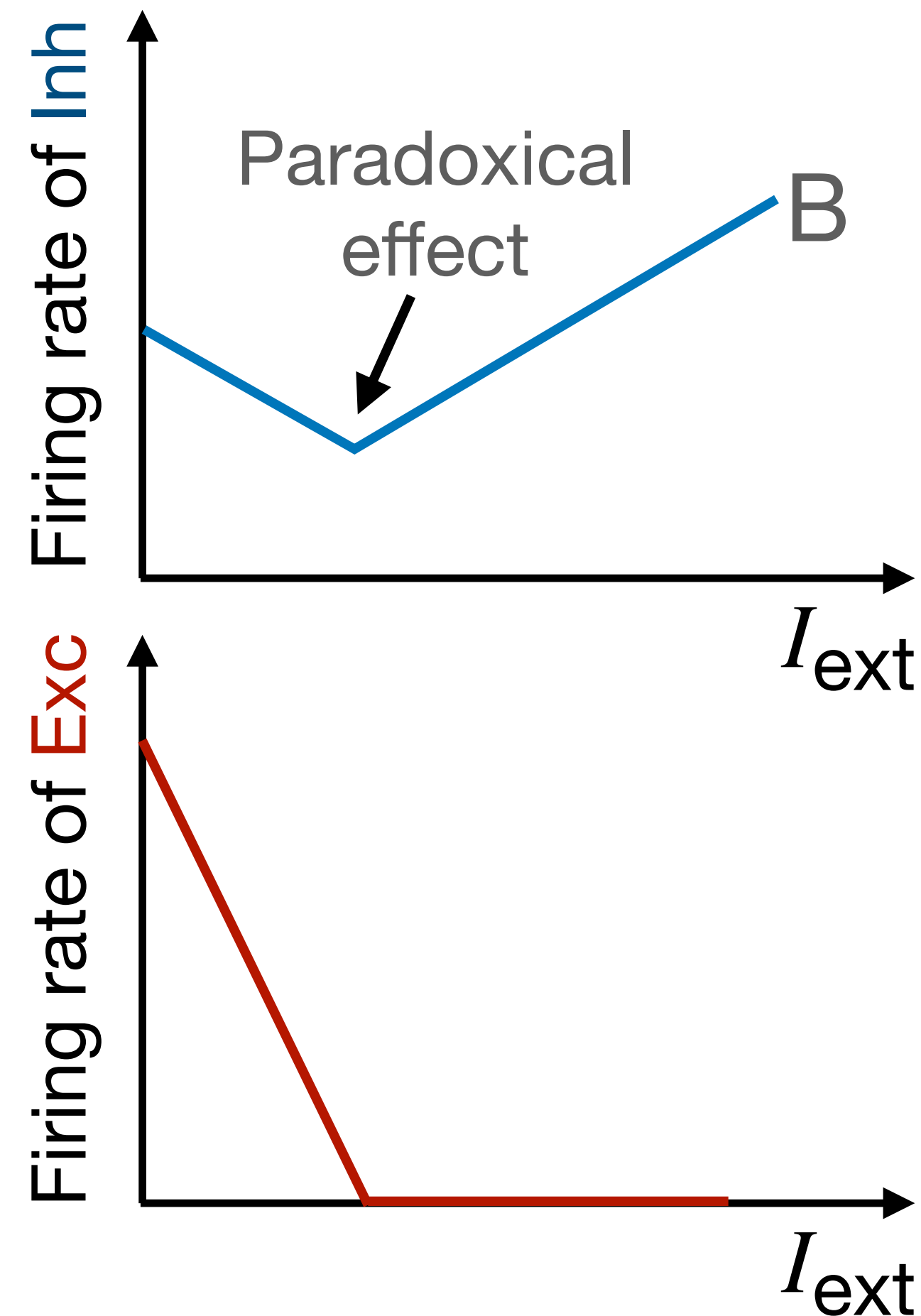
# Paradoxical Effect

*An interesting activity pattern in inhibition-stabilized networks (ISNs)*

Experimentally  
observed!



**Key:** The recurrent excitation was too strong when  $I_{\text{ext}} = 0$ !



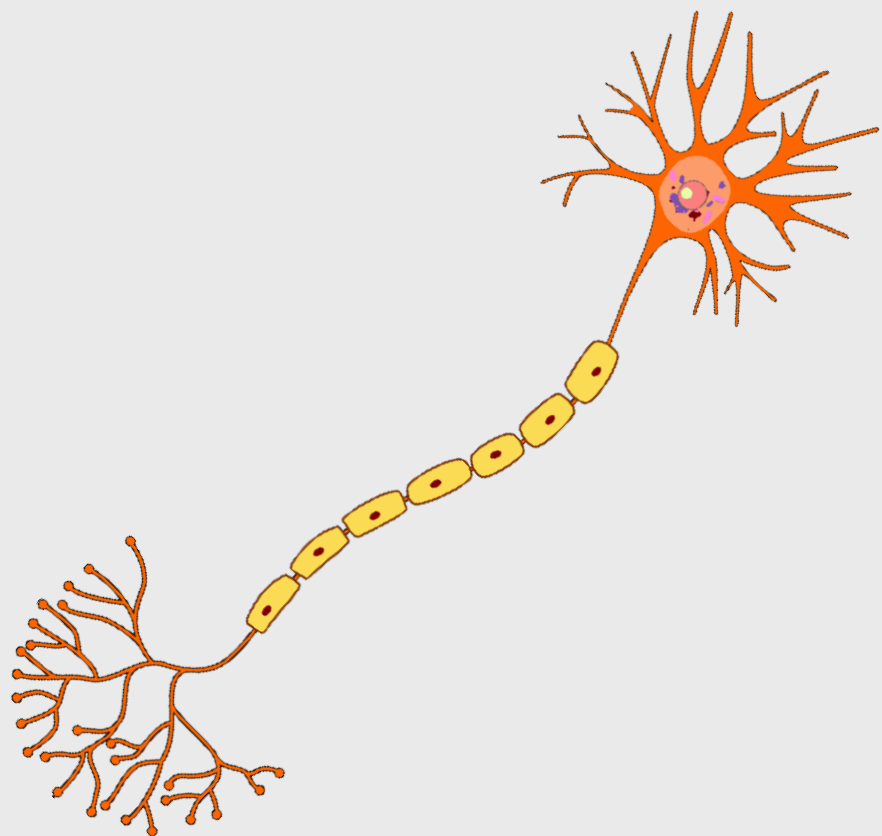
**Q:** What's the role of these motifs?

**Q:** How computation happen in cortex?



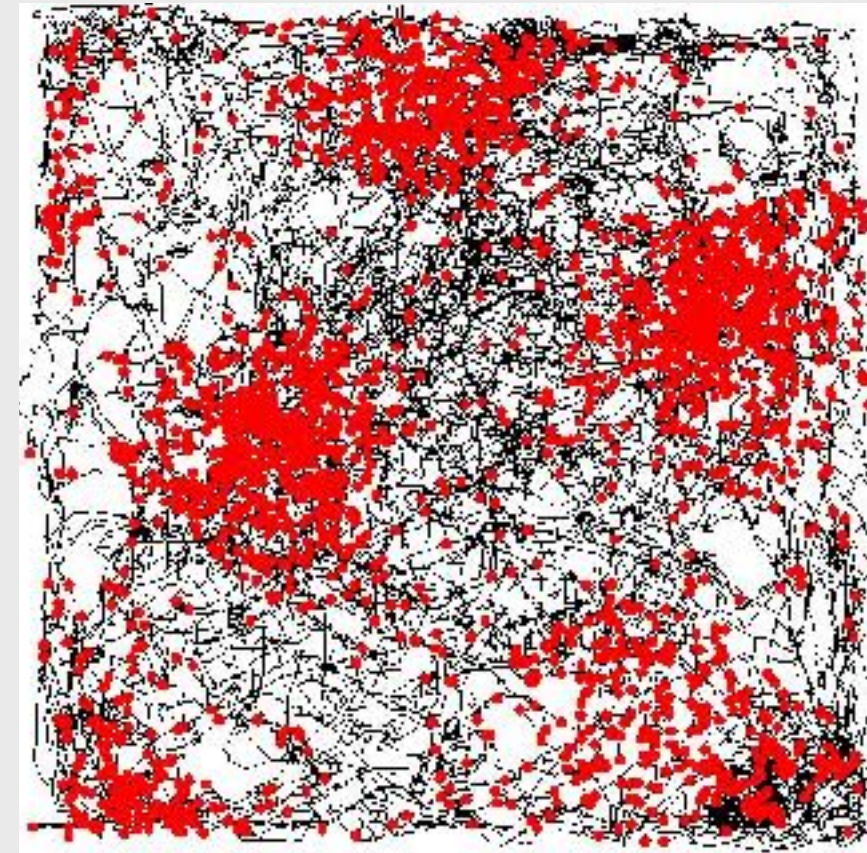
# Beagle Expedition in Neuroscience!?

## Example 1: Single Neuron



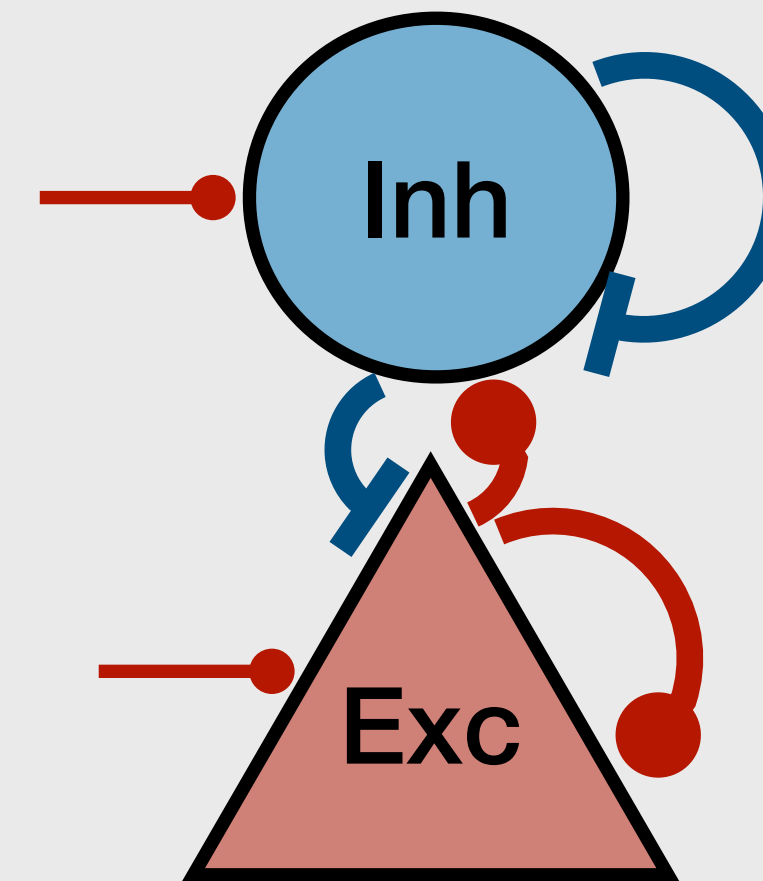
Lowest level

## Example 2: Grid Cells



Near the middle  
of the brain

## Example 3: Inhibition Stabilized Networks



An abstract model

## More!

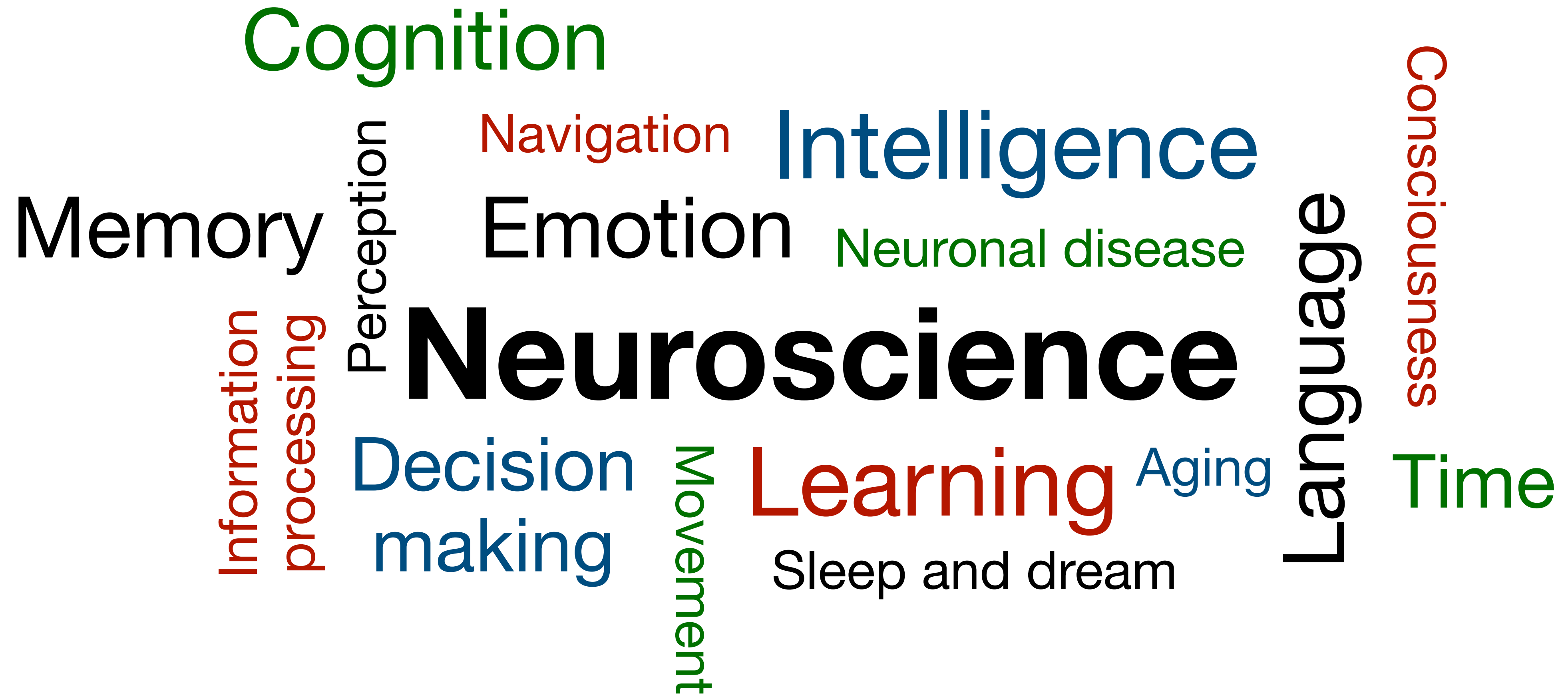


Brabeeba Wang  
(Jan. 20 11am-12pm ET)

Flexible decision  
making in mice



# How Do All the Aspects Harmonize?



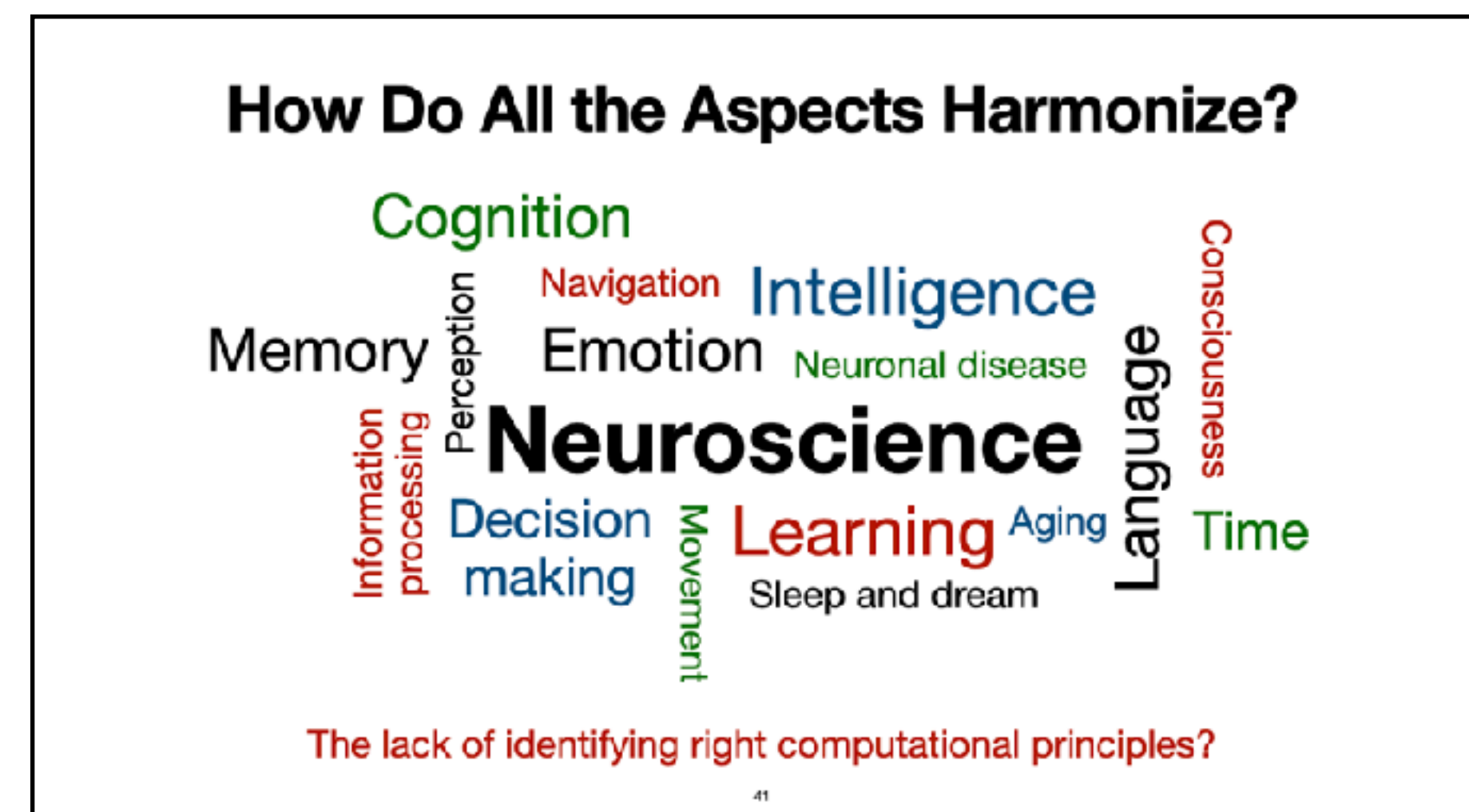
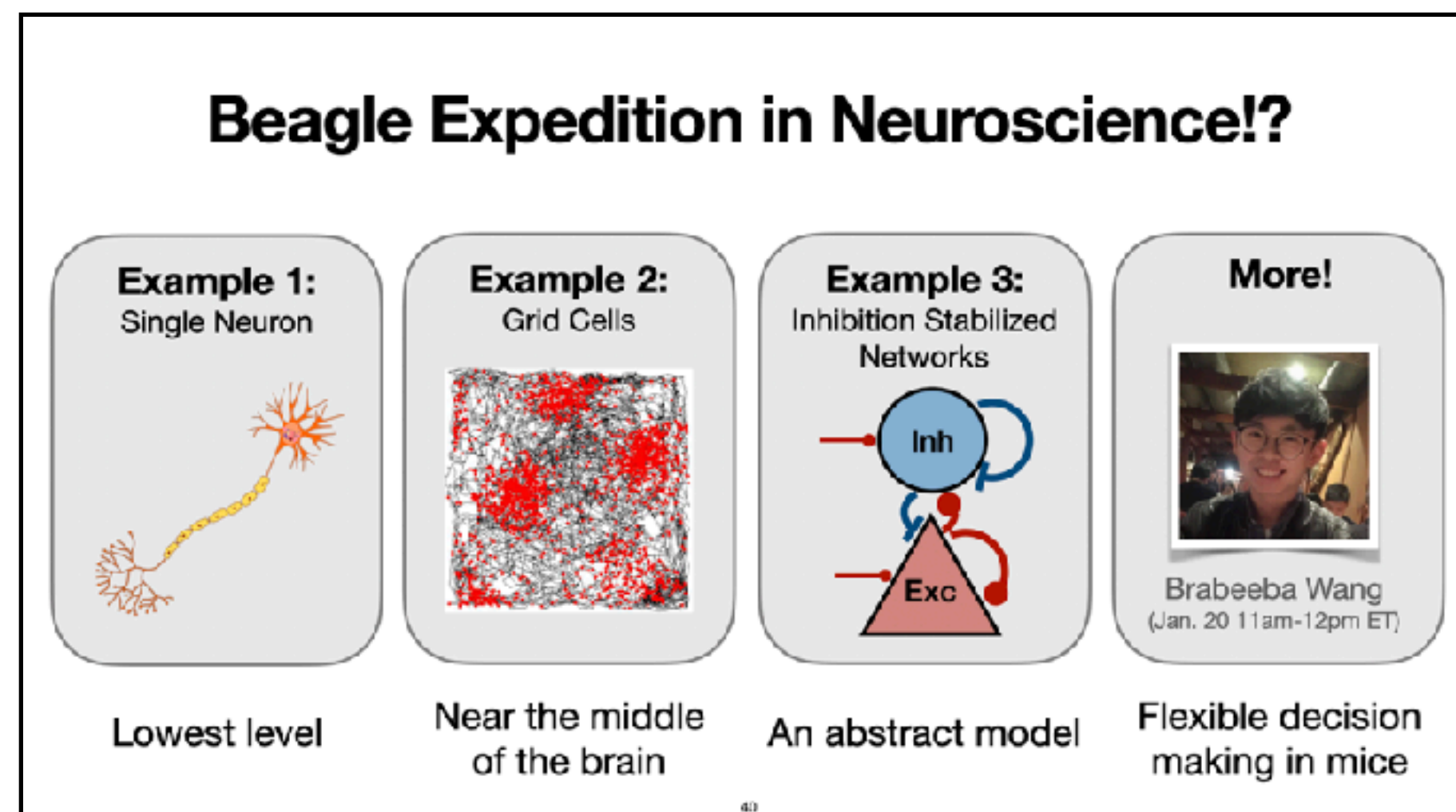
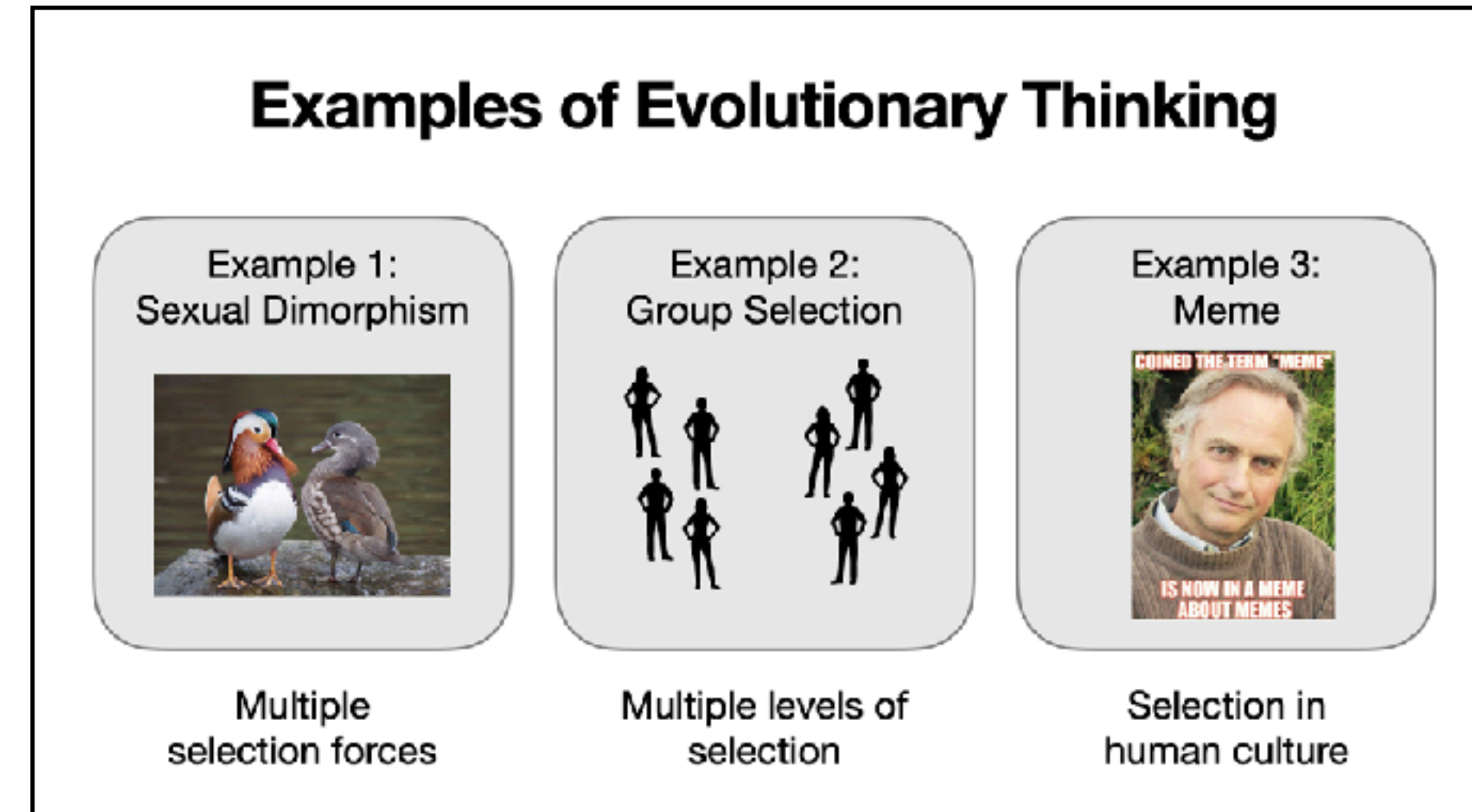
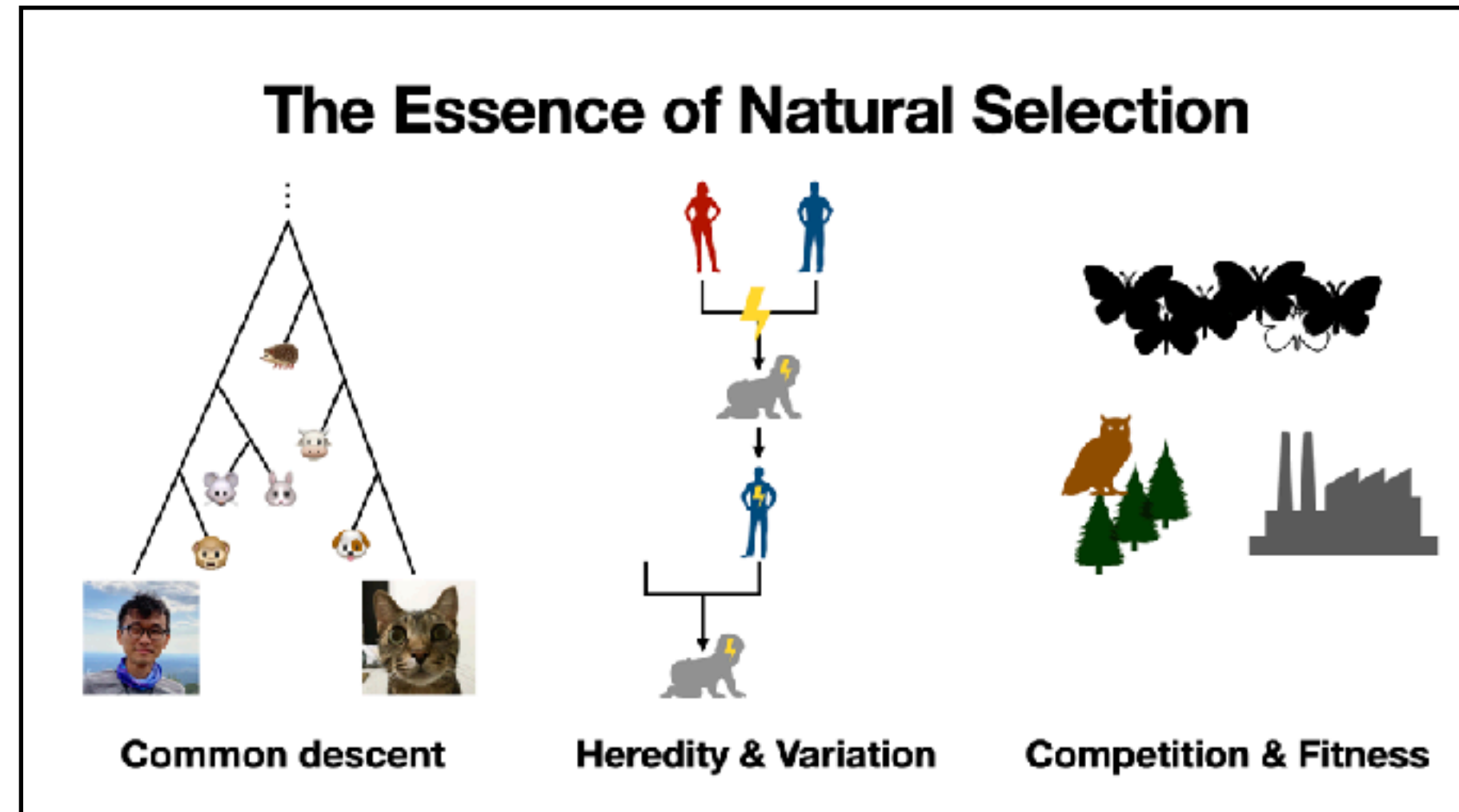
The lack of identifying right computational principles?



# Summary



# Key Concepts



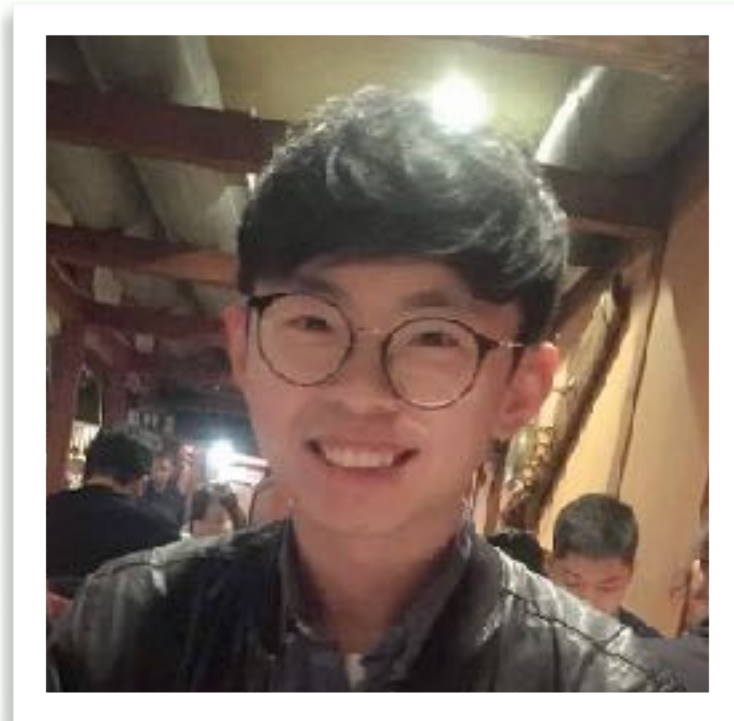


# Guest Speakers for Module III



Angel Hsing-Chi Hwang  
(Jan. 17 11am-12pm ET)

*“Into the Unknown: (De)constructing Creativity in the Age of Human-Machine Partnership”*



Brabeeba Wang  
(Jan. 20 11am-12pm ET)

*“Animal Intelligence: Flexible Computation Under Uncertainty”*



Zhiqian Wang  
(Jan. 19 11am-12pm ET)

*“A Road to Totality: Between Art and Computation”*



# Next



## Lecture III.c

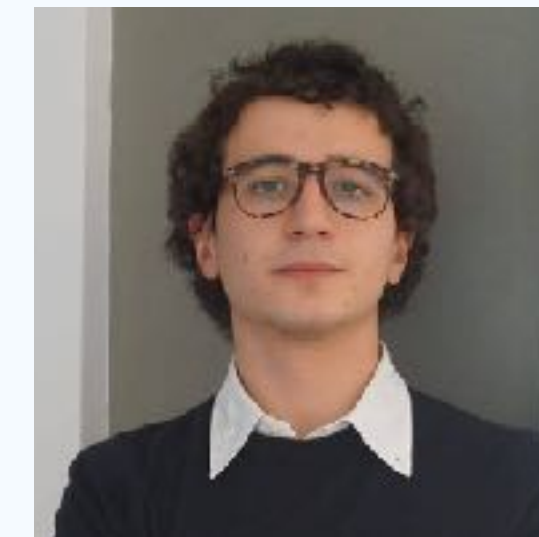
(Jan. 20 10am-10:50am ET)



Salvador

(Jan. 19  
2pm-3pm ET)

*"DNA Computing,  
Cellular Automata, and  
Beyond"*



Simone

(Jan. 14  
2pm-3pm ET)

*"Simulated Annealing"*

Check them out on the calendar!



# References

## Articles:

- Schnitzer, M. Biological computation: Amazing algorithms. Nature 416, 683 (2002), [link](#).
- Chelly Dagdia, Z., Avdeyev, P. & Bayzid, M.S. Biological computation and computational biology: survey, challenges, and discussion. Artif Intell Rev 54, 4169–4235 (2021), [link](#).

## Books:

- Nowak, Martin A. Evolutionary dynamics: exploring the equations of life. Harvard university press, 2006, [link](#).
- Jones, Neil C., Pavel A. Pevzner, and Pavel Pevzner. An introduction to bioinformatics algorithms. MIT press, 2004, [link](#).
- Gillespie, John H. Population genetics: a concise guide. JHU Press, 2004, [link](#).

## Fun reads:

- Stanley, Kenneth O., and Joel Lehman. Why greatness cannot be planned: The myth of the objective. Springer, 2015, [link](#).
- Schrödinger, Erwin. What is life?: With mind and matter and autobiographical sketches. Cambridge university press, 1992, [link](#).
- Mayr, Ernst. This is biology : the science of the living world. Harvard University Press, 2001, [link](#).
- Banatre, Jean-Pierre, et al., eds. Unconventional Programming Paradigms: International Workshop UPP 2004, Le Mont Saint Michel, France, September 15-17, 2004, Revised Selected and Invited Papers. Vol. 3566. Springer Science & Business Media, 2005, [link](#).